



## INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

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<b>(21) International Application Number:</b> PCT/US98/05317  <b>(22) International Filing Date:</b> 18 March 1998 (18.03.98)  <b>(30) Priority Data:</b> 60/044,697                      24 April 1997 (24.04.97)                      US 60/066,135                      19 November 1997 (19.11.97)                      US  <b>(71) Applicant:</b> DOW AGROSCIENCES LLC [US/US]; 9330 Zionsville Road, Indianapolis, IN 46268 (US).	<b>(72) Inventors:</b> PECHACEK, James. T.; 1230 Munsee Court, Indianapolis, IN 46260 (US). TISDELL, Francis, E.; 67 Parkview Road, Carmel, IN 46032 (US). DE VRIES, Donald, H.; 9878 Woodlands Drive, Fishers, IN 46038 (US). SUHR, Robert, G.; 1522 Brunner Drive, Greenfield, IN 46140 (US). JOHNSON, Peter, L.; 8016 North Dartmouth Road, Indianapolis, IN 46260 (US). STOCKDALE, Gary, D.; Apartment 4D, 8722 Knickerbocker Way, Indianapolis, IN 46240 (US). ASH, Mary, L.; 762 Franklin Trace, Zionsville, IN 46077 (US). HAMILTON, Christopher, T.; 8918 Dandy Creek Drive, Indianapolis, IN 46234 (US). HATTON, Christopher, J.; 2525 State Road 38 East, Westfield, IN 46074 (US). JOHNSON, George, W.; 6715 Knollcreek Drive, Indianapolis, IN 46256 (US). YAP, Maurice, Chee, Hoong; 22 Chestnut Court, Zionsville, IN 46077 (US).  <b>(74) Agent:</b> STUART, Donald, R.; Dow AgroSciences LLC, 9330 Zionsville Road, Indianapolis, IN 46268 (US).  <b>(81) Designated States:</b> AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, YU, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).  <b>Published</b> <i>With international search report.</i> <i>Before the expiration of the time limit for amending the</i> <i>claims and to be republished in the event of the receipt of</i> <i>amendments.</i>	
<b>(54) Title:</b> PESTICIDAL 3-(SUBSTITUTED PHENYL)-5-(THIENYL OR FURYL)-1,2,4-TRIAZOLES  <b>(57) Abstract</b>  3-(Substituted phenyl)-5-(thienyl or furyl)-1,2,4-triazole compounds are useful as insecticides and acaricides. New synthetic procedures and intermediates for preparing the compounds, pesticide compositions containing the compounds, and methods of controlling insects and mites using the compounds are also provided.		

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## PESTICIDAL 3-(SUBSTITUTED PHENYL)-5-(THIENYL OR FURYL)-1,2,4-TRIAZOLES

Related Applications

5           This application claims the priority of United States  
Patent Applications Ser. No. 60/044,697, filed April 24, 1997,  
and Ser. No. 60/066,135, filed November 19, 1997.

Field of the Invention

10           This invention provides new compounds that are useful as  
insecticides and acaricides, new synthetic procedures and  
intermediates for preparing the compounds, pesticide  
compositions containing the compounds, and methods of  
controlling insects and mites using the compounds.

Background of the Invention

15           There is an acute need for new insecticides and  
acaricides. Insects and mites are developing resistance to  
the insecticides and acaricides in current use. At least 400  
species of arthropods are resistant to one or more  
insecticides. The development of resistance to some of the  
20   older insecticides, such as DDT, the carbamates, and the  
organophosphates, is well known. But resistance has even  
developed to some of the newer pyrethroid insecticides and  
acaricides. Therefore a need exists for new insecticides and  
acaricides, and particularly for compounds that have new or  
25   atypical modes of action.

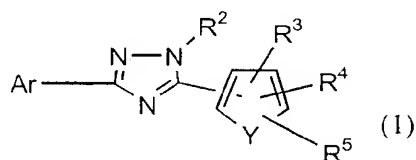
          A number of 3,5-diphenyl-1H-1,2,4-triazole derivatives  
have been described in the literature as having acaricidal  
activity. U.S. 5,482,951; JP 8092224, EP572142, JP 08283261.  
To applicants knowledge, however, none of these compounds has  
30   become a commercial product. Nitro furanyl triazoles are  
described by L.E. Benjamin and H.R. Snyder as antimicrobials  
(*J. Heterocyclic Chem.* 1976, 13, 1115) and by others as

antibacterials (*J. Med. Chem.* **1973**, 16(4), 312-319; *J. Med. Chem.* **1974**, 17(7), 756-758). The present invention provides novel compounds with commercial level activity against mites and insects.

## 5 Summary of the Invention

This invention provides novel substituted thienyl and furanyl triazole derivatives especially useful for the control of insects and mites.

10 More specifically, the invention provides novel insecticidally active compounds of the formula (1)



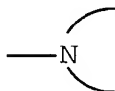
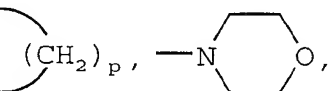
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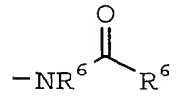
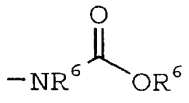
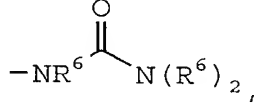
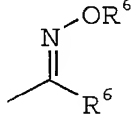
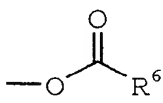
Ar is substituted phenyl;

Y is O or S;

15  $R^2$  is lower alkyl, haloalkyl, lower alkenyl, lower alkynyl, or alkoxyalkyl;

$R^3$  is selected from H, halo, lower alkyl, (C<sub>7</sub>-C<sub>21</sub>) straight or branched chain alkyl, hydroxy, lower alkoxy, haloalkyl, haloalkoxy, alkoxyalkyl, alkoxyalkoxy, lower  
20 alkenyl, lower alkynyl, haloalkenyl, CN, NO<sub>2</sub>, COR<sup>6</sup>, CO<sub>2</sub>R<sup>6</sup>, CON(R<sup>6</sup>)<sub>2</sub>, (C<sub>3</sub>-C<sub>6</sub>) cycloalkyl, S(O)<sub>m</sub>R<sup>6</sup>, -OSO<sub>2</sub>R<sup>6</sup>, SCN, -(CH<sub>2</sub>)<sub>n</sub>R<sup>6</sup>, -CH=CHR<sup>6</sup>, -C≡CR<sup>6</sup>, -(CH<sub>2</sub>)<sub>q</sub>OR<sup>6</sup>, -(CH<sub>2</sub>)<sub>q</sub>SR<sup>6</sup>, -(CH<sub>2</sub>)<sub>q</sub>NR<sup>6</sup>R<sup>6</sup>, -

O(CH<sub>2</sub>)<sub>q</sub>R<sup>6</sup>, -S(CH<sub>2</sub>)<sub>q</sub>R<sup>6</sup>, -NR<sup>6</sup>(CH<sub>2</sub>)<sub>q</sub>R<sup>6</sup>, , ,

, , , , , or

$\text{—S—C(=O)—R}^6$ ,  $\text{—Si(R}^7)_3$  pyridyl, substituted pyridyl, isoxazolyl,  
 substituted isoxazolyl, naphthyl, substituted naphthyl,  
 phenyl, substituted phenyl, thienyl, substituted thienyl,  
 pyrimidyl, substituted pyrimidyl, pyrazolyl, or substituted  
 5 pyrazolyl;

$\text{R}^4$  and  $\text{R}^5$  are independently H, halo, lower alkyl, lower  
 alkoxy, haloalkyl, haloalkoxy, CN,  $\text{CO}_2\text{R}^6$ ,  $\text{CON(R}^6)_2$ , or  $\text{S(O)}_m$   
 alkyl, or

if  $\text{R}^4$  and  $\text{R}^5$  are attached to adjacent carbon atoms, they  
 10 may join to form a 5 or 6 member saturated or unsaturated  
 carbocyclic ring which may be substituted by 1 or 2 halo,  
 lower alkyl, lower alkoxy or haloalkyl groups;

$\text{R}^6$  is H, lower alkyl, haloalkyl, lower alkenyl, lower  
 alkynyl, phenyl, or substituted phenyl;

15  $\text{R}^7$  is lower alkyl;

$m$  is 0, 1, or 2;

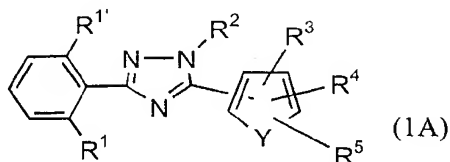
$n$  is 1 or 2;

$p$  is an integer from 2 to 6; and

$q$  is 0 or 1;

20 or a phytologically acceptable acid addition salt thereof.

A preferred group of compounds are those of formula (1A)



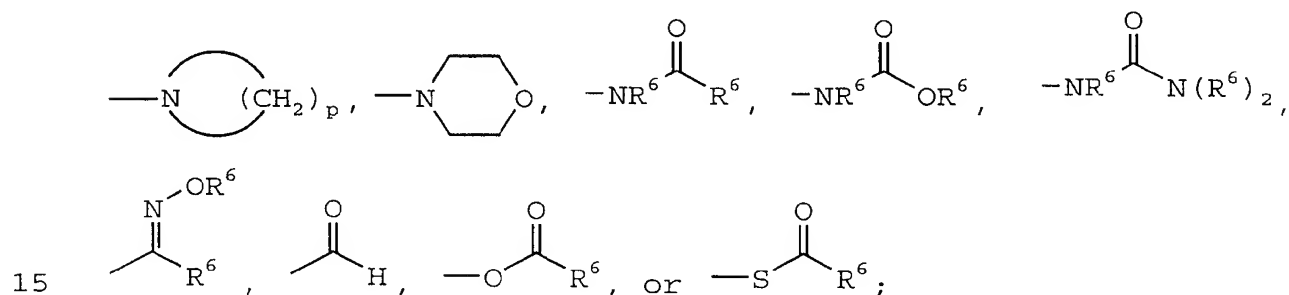
wherein

$R^1$  and  $R^{1'}$  are independently H, Cl, F, methyl, halomethyl, methoxy, or halomethoxy;

Y is O or S;

$R^2$  is lower alkyl, haloalkyl, lower alkenyl, lower alkynyl, or alkoxyalkyl;

$R^3$  is selected from H, halo, lower alkyl,  $(C_7-C_{21})$  straight chain alkyl, hydroxy, lower alkoxy, haloalkyl, haloalkoxy, alkoxyalkyl, alkoxyalkoxy, lower alkenyl, haloalkenyl, CN,  $NO_2$ ,  $CO_2R^6$ ,  $CON(R^6)_2$ ,  $(C_3-C_6)$  cycloalkyl,  $S(O)_mR^6$ , SCN, pyridyl, substituted pyridyl, isoxazolyl, substituted isoxazolyl, naphthyl, substituted naphthyl, phenyl, substituted phenyl,  $-(CH_2)_nR^6$ ,  $-CH=CHR^6$ ,  $-C\equiv CR^6$ ,  $-CH_2OR^6$ ,  $-CH_2SR^6$ ,  $-CH_2NR^6R^6$ ,  $-OCH_2R^6$ ,  $-SCH_2R^6$ ,  $-NR^6CH_2R^6$ ,



$R^4$  and  $R^5$  are independently H, halo, lower alkyl, lower alkoxy, haloalkyl, haloalkoxy, CN,  $CO_2R^6$ ,  $CON(R^6)_2$ , or  $S(O)_m$  alkyl, or

$R^4$  and  $R^5$  form a 5 or 6 member saturated or unsaturated carbocyclic ring which may be substituted by 1 or 2 halo, lower alkyl, lower alkoxy or haloalkyl groups;

$R^6$  is H, lower alkyl, haloalkyl, lower alkenyl, lower alkynyl, phenyl, or substituted phenyl;

m is 0, 1, or 2; and

n is 1 or 2;

p is an integer from 2 to 6;

or a phytologically acceptable acid addition salt thereof.

Preferred compounds of formula (1A) include the following classes:

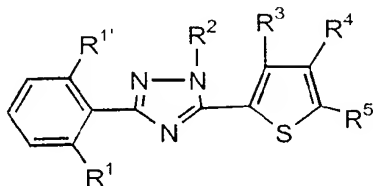
- 5       a) compounds of formula (1A) wherein  $R^{1'}$  is F;
- b) compounds of formula (1A) wherein Y is sulfur;
- c) compounds of formula (1A), and particularly  
compounds of the foregoing classes a) and b), wherein  $R^1$  and  
 $R^{1'}$  are independently Cl or F;
- 10       d) compounds of formula (1A), and particularly  
compounds of the foregoing classes a) through c) wherein  $R^1$   
and  $R^{1'}$  are both F;
- e) compounds of formula (1A), and particularly  
compounds of the foregoing classes a) through c) wherein  $R^1$   
15 and  $R^{1'}$  are both Cl;
- f) compounds of formula (1A), and particularly  
compounds of the foregoing classes a) through c) wherein  $R^1$  is  
Cl and  $R^{1'}$  is F;
- g) compounds of formula (1A), and particularly  
20 compounds of one of the foregoing classes a) through f)  
wherein  $R^2$  is methyl;
- h) compounds of formula (1A), and particularly  
compounds of one of the foregoing classes a) through g)  
wherein  $R^3$ ,  $R^4$ , and  $R^5$  are independently selected from H, halo,  
25 methyl, and methoxy;
- i) compounds of formula (1A), and particularly  
compounds of one of the foregoing classes a) through g)  
wherein  $R^3$ ,  $R^4$ , and  $R^5$  are independently H or halo;

j) compounds of formula (1A), and particularly compounds of one of the foregoing classes a) through g) wherein  $R^3$ ,  $R^4$ , and  $R^5$  are independently H, Cl, or Br;

k) compounds of formula (1A), and particularly compounds of one of the foregoing classes a) through g) wherein  $R^3$ ,  $R^4$ , and  $R^5$  are each halo;

l) compounds of formula (1A), and particularly compounds of one of the foregoing classes a) through g) wherein  $R^3$ ,  $R^4$ , and  $R^5$  are each chloro.

10 A particularly preferred class of compounds includes those of formula (1B)



wherein

$R^1$  and  $R^{1'}$  are independently F or Cl;

15  $R^2$  is lower alkyl, with methyl being most preferred; and  $R^3$ ,  $R^4$  and  $R^5$  are independently H, Cl, or Br.

The invention also provides new processes and intermediates for preparing compounds of formula (1) as well as new compositions and methods of use, which will be  
20 described in detail hereinafter.

#### Detailed Description Of The Invention

Throughout this document, all temperatures are given in degrees Celsius, and all percentages are weight percentages unless otherwise stated.



The term "lower alkyl" refers to (C<sub>1</sub>-C<sub>6</sub>) straight hydrocarbon chains and (C<sub>3</sub>-C<sub>6</sub>) branched and cyclic hydrocarbon groups.

The terms "lower alkenyl" and "lower alkynyl" refer to (C<sub>2</sub>-C<sub>6</sub>) straight hydrocarbon chains and (C<sub>3</sub>-C<sub>6</sub>) branched hydrocarbon groups containing at least one double or triple bond, respectively.

The term "lower alkoxy" refers to -O-lower alkyl.

The terms "halomethyl" and "haloalkyl" refer to methoxy and lower alkyl groups substituted with one or more halo atoms.

The terms "halomethoxy" and "haloalkoxy" refer to methyl and lower alkoxy groups substituted with one or more halo atoms.

The term "alkoxyalkyl" refers to a lower alkyl group substituted with a lower alkoxy group.

The terms "substituted naphthyl", "substituted thienyl", "substituted pyrimidyl", "substituted pyrazolyl", "substituted pyridyl", and "substituted isoxaxolyl" refer to the ring system substituted with one or more groups independently selected from halo, halo (C<sub>1</sub>-C<sub>4</sub>) alkyl, CN, NO<sub>2</sub>, (C<sub>1</sub>-C<sub>4</sub>) alkyl, (C<sub>3</sub>-C<sub>4</sub>) branched alkyl, phenyl, (C<sub>1</sub>-C<sub>4</sub>) alkoxy, or halo (C<sub>1</sub>-C<sub>4</sub>) alkoxy.

The term "substituted phenyl" refers to a phenyl group substituted with one or more groups independently selected from halo, (C<sub>1</sub>-C<sub>10</sub>) alkyl, branched (C<sub>3</sub>-C<sub>6</sub>) alkyl, halo (C<sub>1</sub>-C<sub>7</sub>) alkyl, hydroxy (C<sub>1</sub>-C<sub>7</sub>) alkyl, (C<sub>1</sub>-C<sub>7</sub>) alkoxy, halo (C<sub>1</sub>-C<sub>7</sub>) alkoxy, phenoxy, phenyl, NO<sub>2</sub>, OH, CN, (C<sub>1</sub>-C<sub>4</sub>) alkanoyl, benzoyl, (C<sub>1</sub>-C<sub>4</sub>) alkanoyloxy, (C<sub>1</sub>-C<sub>4</sub>) alkoxycarbonyl, phenoxycarbonyl, or benzoyloxy.

The term "substituted benzenesulfonyl" refers to p-chlorobenzenesulfonyl and p-toluenesulfonyl.

Unless otherwise indicated, when it is stated that a group may be substituted with one or more substituents  
 5 selected from an identified class, it is intended that the substituents may be independently selected from the class.

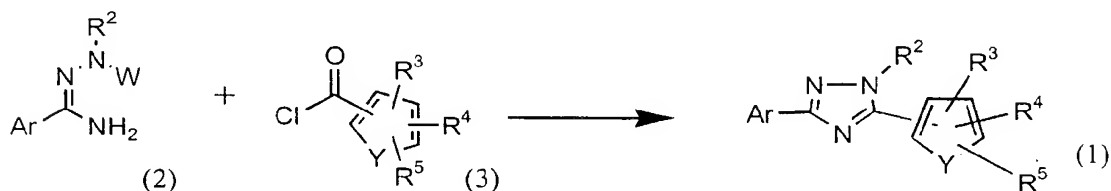
### Synthesis

Compounds of formula (1) can be prepared by the methods described in U.S. Patent Nos. 5,380,944 and 5,284,860  
 10 (Production Methods 1, 2 and 3). Additional methods will be described hereinafter.

For example, compounds of Formula (1) wherein R<sup>1'</sup> is F can be prepared in accordance with the following reaction Scheme I:

15

### Scheme I



wherein Ar, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup> and Y are as defined in formula (1), and W is a conventional amino protecting group. Examples of conventional amino protecting groups include, but are not  
 20 limited to, the carbobenzyloxy group, tertiary alkoxy carbonyl groups, amides, phosphinyl and phosphoryl groups, and sulfenyl and sulfonyl groups. As illustrated in Scheme I, an N-protected amidrazone (2) is reacted with a compound of formula (3) in the presence of acid or base as catalyst.

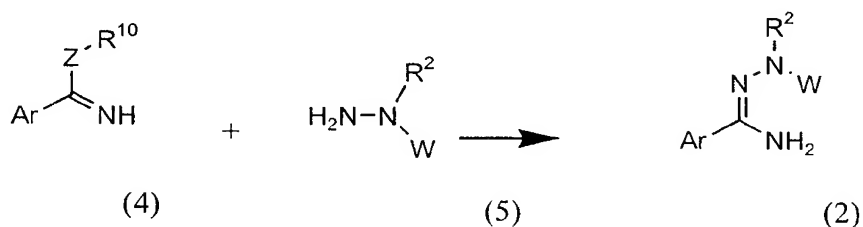
25 Intermediates of formulas (2) and (3) may be obtained by application of well known procedures.

An example of an intermediate of formula (3) is 3,4,5-trichlorothiophene-2-carboxylic acid chloride. The carboxylic

acid can be obtained by the treatment of tetrachlorothiophene with n-butyllithium and subsequent quenching with carbon dioxide as in *J.Organometal.* **1968**, *13*, 419-430. Conversion of carboxylic acid to the acid chloride is carried out by conventional methods, as illustrated hereinafter in Example 2.

Scheme II illustrates preparation of the protected benzamidrazone starting material (2).

Scheme II

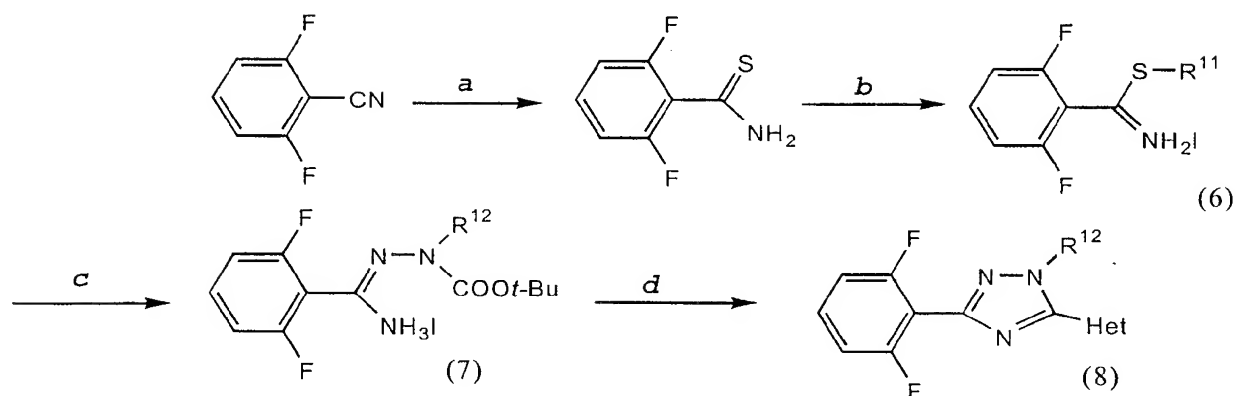


Benzimidate derivative (4), wherein Z is O or S, and R<sup>10</sup> is lower alkyl, is reacted with hydrazine derivative (5), wherein Ar, W, and R<sup>2</sup> are as defined above for Scheme I.

An example of an intermediate of formula (5) is N-methyl-N-t-butylcarboxyhydrazine. Its use in making regiospecific 1-alkyl [1,2,4] triazoles is found in *Chem. Ber.* **1982**, *115*, 2807-2818. The production of benzimide compounds is well known. An example is disclosed in *Synth. Commun.* **1983**, *13*, 753.

Another aspect of the invention is a new method for preparing compounds of formula (1A) wherein R<sup>1</sup> and R<sup>1'</sup> are F, as illustrated in Scheme III:

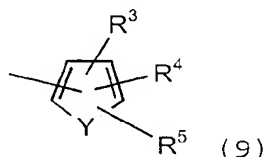
## Scheme III



In Scheme III,  $R^{11}$  is lower alkyl, preferably methyl;

$R^{12}$  is lower alkyl, preferably methyl; and,

5 Het is a thienyl or furyl group of formula (9)



where Y,  $R^3$ ,  $R^4$ , and  $R^5$  are as defined in formula (1). The process illustrated in Scheme III is also applicable to preparation of compounds wherein Het is any of a variety of other heterocyclic groups, for example pyridyl and pyrazolyl.

In another of its aspects, the invention provides novel intermediates of the formulas (6) and (7), as defined above.

As illustrated in step *a* of Scheme III, 2,6-difluorobenzonitrile is reacted with triethylamine, sodium sulfide hydrate, and hydrochloric acid in pyridine at room temperature to give 2,6-difluorobenzenethioamide.

In step *b* of Scheme III the 2,6-difluorobenzenethioamide is reacted with lower alkyl iodide, e.g. iodomethane, in acetone to provide an S-(lower alkyl)thio-2,6-difluorobenzimidinium iodide of formula (6). Acetone is the

preferred solvent, however other polar aprotic solvents such as DMF or THF can be used.

In step *c* of Scheme III the S-(lower alkyl)thio-2,6-difluorobenzimidinium iodide is reacted with an N-t-butoxycarbonyl-N-(lower alkyl)hydrazine to provide the amidrazone of formula (7). The reaction is carried out in methanol or ethanol, preferably methanol, at a temperature of 0°C to the boiling point of the solvent.

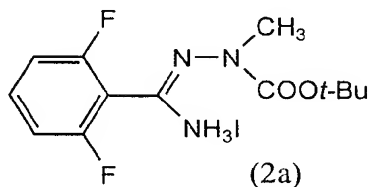
In step *d* of Scheme III, the amidrazone of formula (7) is reacted with a thiophene or furan acid chloride in a nonreactive organic solvent such as benzene, toluene, xylenes, chloroform, dichloromethane, or 1,2-dichloroethane, at a temperature in the range from 0°C to the boiling point of the solvent.

The process of Scheme III uses milder conditions than previously published processes, and therefore allows thermally sensitive heterocycles to be used. Higher yields are also provided.

A detailed illustration of steps *a-c* of Scheme III is given in Example 1 hereinafter. Detailed illustrations for step *d* are given in Examples 2-4 hereinafter.

#### Example 1

The following steps illustrate preparation of the amidrazone of formula (2a)



#### A. 2,6-difluorobenzenethioamide.

Into a 3 L three necked round bottom flask equipped with a mechanical stirrer, dry ice condenser, dropping funnel, and

outlet to a trap filled with bleach was added pyridine (550 mL), 2,6-difluorobenzonitrile (208 g, 1.50 mol), triethylamine (202 g, 279 mL, 2.0 mol), and sodium sulfide hydrate (521 g, 2.17 mol-broken into pieces small enough to fit into the flask). The temperature of the stirred mixture was lowered to approximately 5 °C and to the slurry was added dropwise concentrated hydrochloric acid (143 g, 288 mL, 3.99 mol). An exotherm was noted and the rate of addition of the hydrochloric acid was such that the temperature of the reaction mixture did not exceed 25 °C for a total addition time of 75 min. The cooling bath was removed and the slurry was allowed to warm to RT and to stir over night. The mixture was poured into water (2 L) and was extracted with ether (3 X 500 mL). The ether layer was washed with dilute sulfuric acid, water, brine, dried (MgSO<sub>4</sub>), and the solvent removed in vacuo to give 232 grams of crude product. The starting material was removed from the product via kugelrohr distillation to give 197 g (76%) of 2,6-difluorobenzenethioamide. This material was used without further purification.

B. S-methylthio-2,6-difluorobenzamidinium iodide

Into a 3 L three necked flask equipped with a mechanical stirrer and dropping funnel was added acetone (1150 mL) and 2,6-difluorobenzenethioamide (197 g, 1.14 mol). The temperature of the stirred solution was lowered to approximately 5 °C and iodomethane (161 g, 70.6 mL, 1.14 mol) was added dropwise. The ice bath was removed and the slurry was allowed to stir over night. The resulting yellow solids were removed via filtration and washed with ether to obtain 223 grams. An additional portion of material was obtained from the filtrate by removal of the solvent in vacuo. Ether was added to the residue and the resulting solids removed via filtration to obtain an additional 57 grams of material. The

combined solids totaled 280 g (77.9% yield) of S-methylthio-2,6-difluoro-benzimidinium iodide: mp 168-169 °C; <sup>1</sup>H NMR (DMSO-d<sub>6</sub>) δ 7.7 (m, 1H), 7.4 (m, 2H), 2.7 (s, 3H).

C. N-tert-butoxycarbonyl-N-methylhydrazine

5        Into a 1 L three necked round bottom flask equipped with a mechanical stirrer and dropping funnel was added methylhydrazine (42.2 g, 0.916 mol) and THF (100 mL). The temperature of the mixture was cooled to 5 °C and a solution of di-tert-butyl dicarbonate (100 g, 0.458 mol) dissolved in  
10 THF (150 mL) was added dropwise. The cooling bath was removed and the mixture was stirred at RT overnight. The liquid was decanted from a gummy precipitate and the solvent removed *in vacuo* to give approximately 70 grams of a clear liquid. The gummy precipitate was partitioned between methylene chloride  
15 and water. The methylene chloride was washed with brine, dried (Na<sub>2</sub>SO<sub>4</sub>) and the solvent removed *in vacuo*. The resulting residue was combined with that from the previous evaporation and distilled at approximately 20 mm Hg (bp 77-78 °C) to give 40.2 g (60% yield) of N-tert-butoxycarbonyl-N-methylhydrazine:  
20 <sup>1</sup>H NMR (CDCl<sub>3</sub>) δ 4.1 (s, b, 2H), 3.05 (s, 3H), 1.5 (s, 9H).

D. Amidrazone of formula (2a)

Into a 1 L round bottom flask equipped with a mechanical stirrer, dropping funnel, and outlet to a trap filled with bleach, was added S-methyl-2,6-difluorobenzimidinium iodide  
25 (63.8 g, 0.202 mol) and methanol (180 mL). To the stirred solution was added dropwise N-tert-butoxycarbonyl-N-methylhydrazine (29.6 g, 0.202 mol). The solution was allowed to stir overnight and the methanol was removed *in vacuo*. The residue was triturated with ether and the solids removed via  
30 filtration to give 66.3 grams (79.0% yield) of the amidrazone of formula (2a): mp 172-173 °C (dec); <sup>1</sup>H NMR (DMSO-d<sub>6</sub>) δ 12.3

(s, b, 1H), 10.4 (d, b, 2H), 7.9 (m, 1H), 7.4 (m, 2H), 3.1 (s, 3H), 1.5 (s, 9H).

### Example 2

5                    3-(2,6-difluorophenyl)-5-(3-n-hexylthien-2-yl)-1-methyl[1,2,4]triazole (Compound 35)

To a mixture of 3-n-hexyl-2-thiophene carboxylic acid (0.5 g, 2.4 mmol) in 25 mL of dry 1,2-dichloroethane was added 2.0 mL of thionyl chloride and a drop of dimethylformamide. The mixture was refluxed for four hours. After cooling, the  
10 reaction mixture was evaporated in vacuo and the residue was combined with 25 mL of dry toluene and 1.24 g (3.0 mmol) of the amidrazone of formula (2a) (Example 1). The mixture was refluxed overnight, cooled, and then partitioned between brine and ether. The organic phase was dried (MgSO<sub>4</sub>), the solvent  
15 evaporated, and the residue was chromatographed on silica gel using ethyl acetate/hexanes as the eluant; 5:95 to 20:80. The product fractions were collected and evaporated to give 0.66 g (76% yield) of the title product as an oil. <sup>1</sup>H NMR δ 7.26-7.46 (m, 2H), 6.97-7.06 (m, 2H), 3.97 (s, 3H) 2.71 (m, 2H), 1.23  
20 (m, 6H), 0.84 (m, 3H); calcd. for C<sub>19</sub>H<sub>21</sub>F<sub>2</sub>N<sub>3</sub>S: C, 63.14; H, 5.86; N, 11.63; S, 8.87; Found: C, 62.90 ; H, 5.79; N, 11.70; S, 8.77.

### Example 3

25                    3-(2,6-difluorophenyl)-5-(4-p-chlorobenzenesulfonyl-3-methylthien-2-yl)-1-methyl[1,2,4]triazole (Compound 12)

To a solution of 0.9 g (2.2 mmol) of the amidrazone of formula (2a) (Example 1) and 0.75 g (2.2 mmol) of the thiophene acid chloride, 4-(p-chlorobenzenesulfonyl)-3-methylthiophene-2-carbonyl chloride, in 100 mL of toluene was added 1  
30 equivalent (0.4 g, 2.2 mmol) p-toluenesulfonic acid monohydrate with stirring. The mixture was refluxed overnight



using a Dean-Stark trap to remove water. The solution was cooled and the solvent removed *in vacuo*. The residue was dissolved in 100 mL of dichloromethane and was washed with 2N NaOH and then with water. The organic phase was dried (MgSO<sub>4</sub>) and the solvent evaporated. The residue was recrystallized from ethyl acetate / hexanes to give 0.6 g of the product as off-white crystals. mp 183-185 °C. 59.4% yield. <sup>1</sup>H NMR δ 8.4 (s, 1H), 7.8 (d, 2H), 7.4 (d, 2H) 7.3 (m, 1H), 7.0 (m, 2H), 3.98 (s, 3H), 2.25 (s, 3H); calcd. for C<sub>20</sub>H<sub>14</sub>F<sub>2</sub>ClN<sub>3</sub>O<sub>2</sub>S<sub>2</sub>: C, 51.55; H, 3.01; N, 9.02; Found: C, 51.76 ; H, 3.26; N, 8.92.

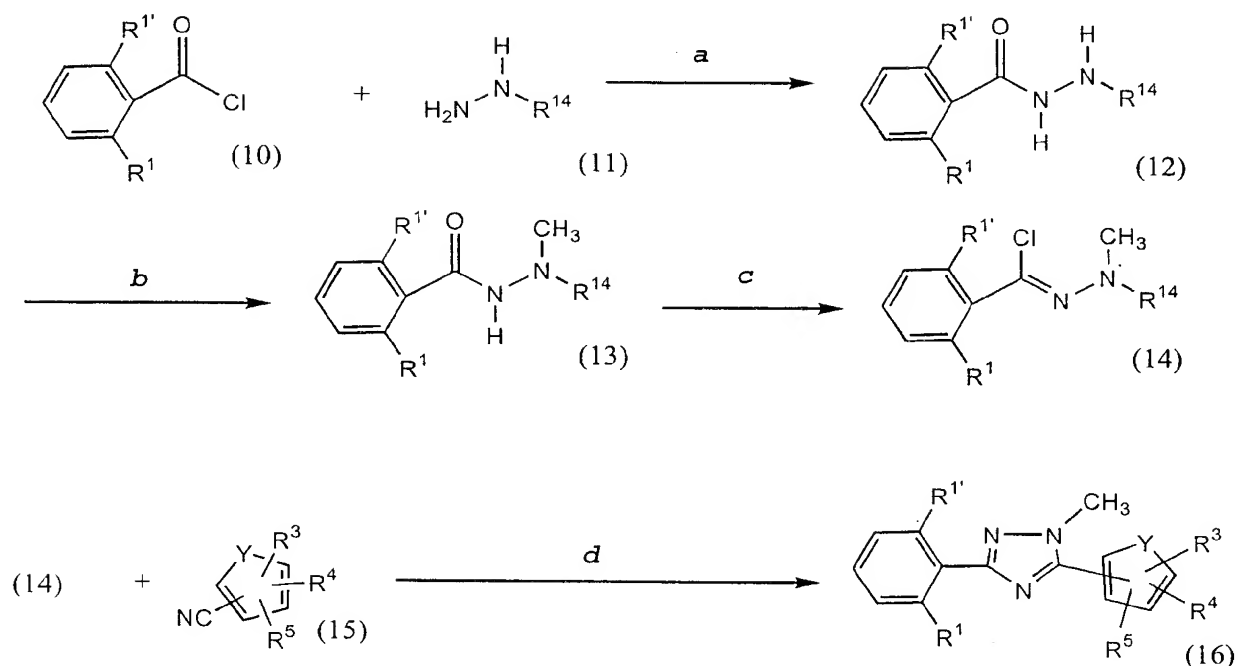
#### Example 4

##### 3-(2,6-difluorophenyl)-5-(5-p-chlorophenyl)furan-2-yl)-1-methyl[1,2,4]triazole (Compound 13)

To a solution of 1.0 g (2.4 mmol) of the amidrazone of formula (2a) (Example 1) and 0.61 g (2.4 mmol) of 5-[p-chlorophenyl]-furan-2-carbonyl chloride in 100 mL of toluene was added 1 equivalent (0.43 g, 2.4 mmol) p-toluenesulfonic acid monohydrate with stirring. The mixture was refluxed overnight using a Dean-Stark trap to remove water. The solution was cooled and the solvent removed *in vacuo*. The residue was dissolved in 100 mL of dichloromethane and was washed with 2N NaOH and then with water. The organic phase was dried (MgSO<sub>4</sub>) and the solvent evaporated. The residue was recrystallized from ethyl acetate/hexanes to give 0.4 g of the product as off white crystals mp 171-173 °C. 44.9% yield; calcd. for C<sub>19</sub>H<sub>12</sub>F<sub>2</sub>ClN<sub>3</sub>O: C, 61.37; H, 3.23; N, 11.30; Found: C, 61.34 ; H, 3.19; N, 11.06.

The invention also provides a new method for preparing compounds of formula (1A) as illustrated in Scheme IV:

Scheme IV



wherein  $R^1$ ,  $R^{1'}$ , Y,  $R^3$ ,  $R^4$ , and  $R^5$  are as defined in formula (1) and  $R^{14}$  is methanesulfonyl, benzenesulfonyl, or substituted benzenesulfonyl.

In step *a*, 2,6-substitutedbenzoylchloride (10) is reacted with a hydrazide of formula (11) (wherein  $R^{14}$  is methanesulfonyl, benzenesulfonyl, or substituted benzenesulfonyl, i.e. p-Cl or p-CH<sub>3</sub> benzenesulfonyl) and triethylamine in THF to give the substituted benzhydrazone of formula (12).

In step *b*, the substituted benzhydrazone of formula (12) is reacted first with sodium hydride in N,N-dimethylformamide and then with iodomethane to produce the substituted hydrazine of formula (13).

In step *c* the substituted hydrazine of formula (13) is chlorinated using, e.g. PCl<sub>5</sub> to produce substituted benzylhydrazonoyl chloride of formula (14). The reaction is

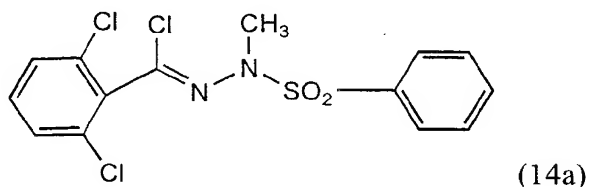
carried out in a non-reactive organic solvent such as dichloroethane.

In step *d* the substituted benzylhydrazonoyl chloride of formula (14) is reacted with a mixture of aluminum chloride and cyanothiophene or cyanofuran of formula (15) in a solvent, e.g. o-dichlorobenzene, to produce a 3-(2,6-substituted phenyl)-5-(substituted thienyl)-1-methyl[1,2,4]triazole of formula (16).

A detailed illustration of steps *a-c* of Scheme IV is given in Example 5 hereinafter. Detailed illustrations of step *d* are given in Examples 6-9, 11, 13, and 15 hereinafter.

#### Example 5

The following steps show preparation of the benzhydrazoneoyl chloride of formula (14a)



#### A. 1-Benzenesulfonyl-2-(2,6-dichloro)benzhydrazone

Into a 1 L three necked round bottom flask equipped with a condenser, mechanical stirrer, thermometer, under an atmosphere of nitrogen, was added THF (500 mL), benzenesulfonyl hydrazide (41.1 g, 0.238 mol), and triethylamine (24.1 g, 33.2 mL, 0.238 mol). The resulting solution was cooled to -5 °C and 2,6-dichlorobenzoyl chloride (50.0 g, 34.2 mL, 0.238 mol) was added dropwise over a 55-min period and the temperature did not rise above 0°C. The reaction mixture was allowed to stir for 1 hr and then the cooling bath was removed and the reaction mixture was then stirred 21 hr at room temperature with monitoring via TLC and HPLC. Most of the solvent was removed *in vacuo* and the

residue partitioned between methylene chloride (1000 mL) and water (2 X 200 mL). The organic layer was washed with saturated brine (250 mL), dried ( $\text{Na}_2\text{SO}_4$ ), and the solvent removed *in vacuo* to give a white solid. The white solids were slurried in ether and were removed via filtration and dried *in vacuo* overnight to give 74.3 g (90.5 % yield of 1-benzenesulfonyl-2-(2,6-dichlorobenz)hydrazone: mp 180-181 °C; TIC mass 345/347/349;  $^1\text{H}$  NMR ( $\text{CDCl}_3$ )  $\delta$  8.05 (m, 2H), 7.9 (m, 1H), 7.5-7.7 (m, 4H), 7.3 (m, 3H).

10 B. 1-(2,6-dichloro)benzoyl-2-methyl-2-benzenesulfonyl hydrazine

In a 1 L three necked round bottom flask equipped with a mechanical stirrer, thermometer, and dropping funnel under an atmosphere of nitrogen a suspension of sodium hydride (8.49 g of 60% dispersion, 0.212 mol) was washed with hexanes (3 portions) and most of the hexane from the final wash was removed via suction. N,N-dimethylformamide (200 mL) was added and the temperature of the slurry lowered to -5 °C. A solution of N-benzenesulfonyl-2,6-dichloro-benzhydrazone (73.3 g, 0.212 mol, in 300 mL of N,N-dimethylformamide) was added dropwise over a 120 min period at a rate such that the temperature did not rise above 3 °C and the rate of hydrogen evolution was maintained at a manageable rate. As the addition proceeded the mixture turned lemon yellow and thickened, but when the addition was completed the mixture was clear and easily stirred. The resulting mixture was stirred at 0 °C for 1 hr and the cooling bath removed and stirred for an additional 1 hr (temperature rose to 15 °C). The mixture was then cooled to -5 °C and iodomethane (30.0 g, 13.2 mL) was added dropwise at a rate such that the temperature did not rise above 0 °C. The cooling bath was removed and the reaction mixture was allowed to warm to room temperature and

to stir for 2 hr. The reaction mixture was diluted with brine (300 mL) and extracted with ethyl acetate. The organic layer was washed with saturated brine, dried ( $\text{Na}_2\text{SO}_4$ ), and the solvent removed *in vacuo* to give crude material contaminated with N,N-dimethylformamide which was removed via vacuum pump. The crude mixture was crystallized in hexanes/ethyl acetate, the solids removed via filtration, and dried *in vacuo* to give 40.2 g. Solvent was removed from the filtrate to obtain an additional 21.4 g for a total of (80.9 % yield) of 1-(2,6-dichlorobenzoyl)-2-methyl-2-benzenesulfonyl hydrazine: mp 177-178 °C;  $^1\text{H}$  NMR ( $\text{CDCl}_3$ )  $\delta$  8.0 (m, 2H), 7.4-7.8 (m, 4H), 7.2 (m, 3H), 3.4 and 3.05 (two singlets combined for 3H).

C. N-(Benzenesulfonyl)-N-methyl-(2,6-dichlorobenz)hydrazonoyl chloride

Into a 1 L three necked round bottom flask equipped with a magnetic stirrer and condenser under an atmosphere of nitrogen was added N-benzenesulfonyl-N-methyl-2,6-dichlorobenzoyl hydrazine (35.9 g, 0.10 mol), 1,2-dichloroethane (500 mL), and phosphorus pentachloride (31.2 g, 0.15 mol). The temperature of the mixture was raised to the point of reflux and was allowed to stir for 30 min. The solvent was removed *in vacuo* and the residue dissolved in methylene chloride and carefully diluted with water. The organic phase was washed with water, brine, dried ( $\text{Na}_2\text{SO}_4$ ) and the solvent removed *in vacuo* to give an oil which solidified when triturated with ethyl acetate/hexanes mixture to give 36.1 g (95.7 % yield) N-benzenesulfonyl-N-methyl-2,6-dichlorobenzhydrazonoyl chloride as a white crystalline solid: mp 103-104 °C;  $^1\text{H}$  NMR ( $\text{CDCl}_3$ )  $\delta$  7.9 (m, 2H), 7.4-7.7 (m, 3H), 7.2-7.4 (m, 3H), 3.1 (s, 3H).

N-Benzenesulfonyl-N-methyl-2-chloro-6-fluorobenzhydrazonoyl chloride was also made using the same procedure.

#### Example 6

5     3-(2-chloro-6-fluorophenyl)-5-(3,4,5-trichlorothiophen-2-yl)-1-methyl[1,2,4]triazole (Compound 2)

To a mixture of N-methyl-N-benzenesulfonyl-2-chloro-6-fluorobenzhydrazonoyl chloride (47.5 g, 0.131 mol) and 2-cyano-3,4,5-trichlorothiophene (26.5 g 0.125 mol in o-dichlorobenzene (18 mL), was added aluminum chloride (11.1 g, 0.083 mol). The mixture was lowered into a preheated oil bath maintained at 140-150 °C. The mixture was allowed to remain in the hot oil bath 40 min and was then removed from the bath and stirred for 2 hr as it gradually cooled. The reaction mixture was poured into 2N sodium hydroxide (300 mL-enough to dissolve the aluminum salts) and extracted with methylene chloride (3 X 250 mL). The organic phases were combined, washed with brine, dried (MgSO<sub>4</sub>), and the solvent removed in vacuo to give the crude product. The residue was placed on silica gel (150 g) and then chromatographed using ethyl acetate/hexanes as eluant to obtain 39.7 g (55.3% yield) of the title compound; 99.4% pure by GC analysis; mp 129-130 °C; <sup>1</sup>H NMR (CDCl<sub>3</sub>) δ 7.3 (m, 2H), 7.1 (m, 1H), 4.0 (s, 3H); Anal. for C<sub>13</sub>H<sub>6</sub>Cl<sub>4</sub>FN<sub>3</sub>S: Calcd. C, 39.32; H, 1.52; N, 10.58. Found C, 39.08; H, 1.30; N, 10.34.

#### Example 7

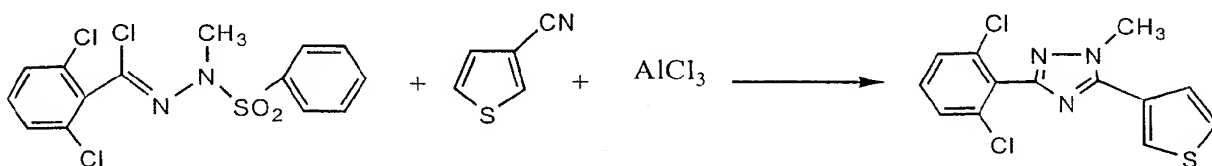
3-(2,6-dichlorophenyl)-5-(3,4,5-trichlorothiophen-2-yl)-1-methyl[1,2,4]triazole (Compound 45)

Into a 500 mL round bottom flask equipped with a mechanical stirrer and condenser, under an atmosphere of nitrogen, was added o-dichlorobenzene (150 mL), 2-cyano-3,4,5-

trichlorothiophene (23.6 g, 111 mmol), and aluminum chloride (14.8 g, 111 mmol). The stirred reaction mixture was plunged into an oil bath at 145-150 °C. When the internal temperature of the mixture reached 133 °C N-(benzenesulfonyl)-N-methyl-  
5 (2,6-dichlorobenz)hydrazonoyl chloride (35.0 g, 92.6 mmol) was added in 5 gram portions over 40 min. The temperature rose to 141 °C. The reaction mixture was allowed to stir from the initial addition of the N-(benzenesulfonyl)-N-methyl-(2,6-dichlorobenz)hydrazonoyl chloride 1 hr and 40 min with  
10 monitoring via HPLC and GC. The oil bath was dropped down from the reactor and the mixture was allowed to cool to approximately 90 °C and was added to stirred 2N caustic (350 mL). The mixture stirred for 5 min, was checked to insure that the pH was basic, and was diluted with methylene chloride  
15 (800 mL). The methylene chloride layer was washed with water (250 mL), brine (250 mL), dried (Na<sub>2</sub>SO<sub>4</sub>), the solvent removed *in vacuo* and the o-dichlorobenzene and benzenesulfonyl chloride were removed via kugelrohr distillation. The straw-colored solids were recrystallized from ethyl acetate to give  
20 31.3 g (81.7% yield) of the title compound; mp 146-147°C; <sup>1</sup>H NMR (CDCl<sub>3</sub>) δ 7.2-7.5 (m, 3H), 4.0 (s, 3H).

## Example 8

3-(2,6-dichlorophenyl)-5-(thien-3-yl)-1-methyl[1,2,4]triazole  
25 (Compound 48)



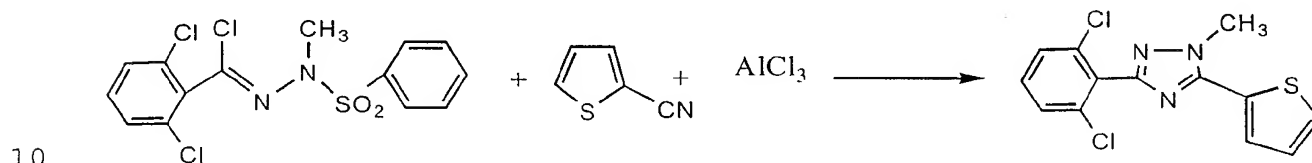
N-(Benzenesulfonyl)-N-methyl-(2,6-dichlorobenz)hydrazonoyl chloride (1.13g, 3mmol), 3-cyanothiophene (0.327g, 6 mmol), and aluminum chloride (0.40 g, 3 mmol) were combined and

heated at 135-140°C for approximately 14 hours, and then stirred at room temperature for approximately 48 hours. The mixture was then diluted with dichloromethane, washed with water, washed with brine, and dried over magnesium sulfate.

5 Chromatography produced 290 mg of the title product as a clear oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>) δ 7.27-7.85 (m, 5H), 4.15 (s, 3H).

Example 9

3-(2,6-dichlorophenyl)-5-(thien-2-yl)-1-methyl[1,2,4]triazole  
(Compound 50)

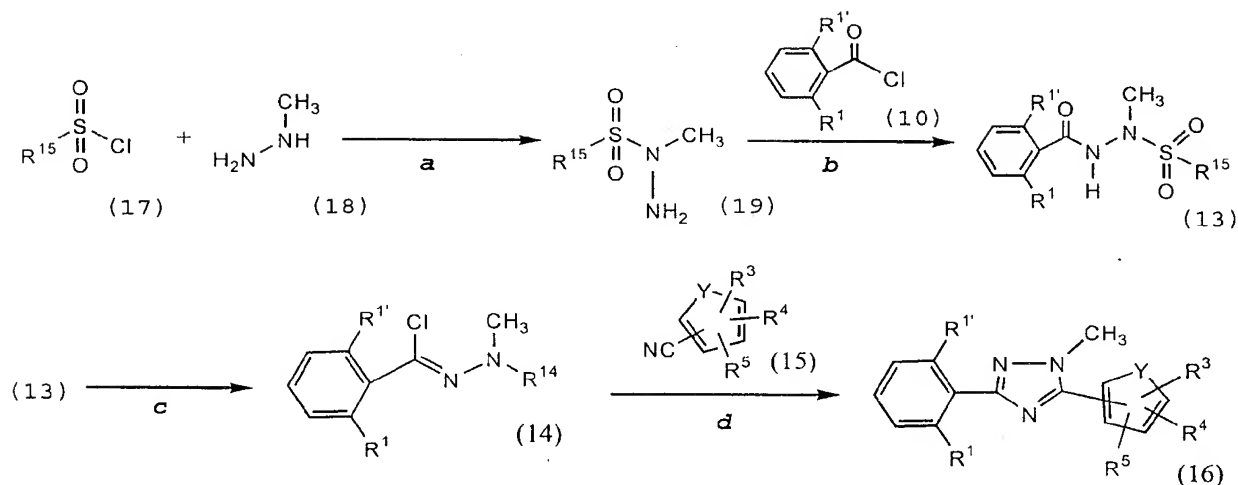


N-Benzenesulfonyl-N-methyl-2,6-dichlorobenzhyrazonoyl chloride (0.42 g, 1.1 mmol), 2-cyanothiophene (0.24g, 2.2 mmol), and aluminum chloride (0.147 g, 1.1 mmol) were combined and heated at 138°C for approximately eight hours. The mixture was then  
15 poured into an ice bath of 1 M NaOH, and stirred one hour, extracted into dichloromethane, washed with brine, dried over magnesium sulfate and concentrated to 230 mg of an oil. Chromatography (SiO<sub>2</sub>, 10% EtOAc-Hex) afforded the product (100 mg) as a white solid. MP 161-163°C.

20 The invention also provides the process for making compounds of formula (16) shown in Scheme V:



## Scheme V



wherein  $R^{15}$  is methyl, phenyl, p-chlorophenyl, or p-tolyl,  $R^{14}$  is methanesulfonyl, benzenesulfonyl, or substituted  
 5 benzenesulfonyl, and  $R^1$ ,  $R^{1'}$ ,  $R^3$ ,  $R^4$ , and  $R^5$  are as defined  
 above for formula (1).

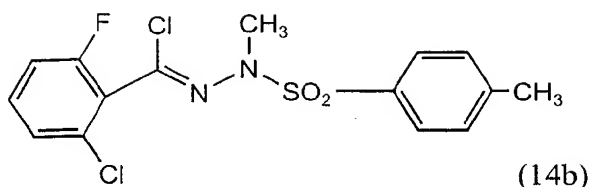
In step **a** of Scheme V, the sulfonyl chloride of formula  
 (17) is reacted with methyl hydrazine in a nonreactive organic  
 solvent such as THF in the presence of triethylamine. The  
 10 reaction can be carried out at room temperature as is  
 typically complete in four hours.

In step **b** of Scheme V, the sulfonyl hydrazine of formula  
 (19) is reacted with the 2,6-substituted benzoyl chloride of  
 formula (10) to provide the intermediate of formula (13).

15 Steps **c** and **d** of Scheme V are the same as steps **c** and **d**  
 of Scheme IV.

## Example 10

This Example shows preparation of the benzhydrazonoyl  
 chloride of formula (14b):



A. 2-methyl-2-(4-methylphenyl)sulfonyl hydrazine

Into a 100 mL three necked round bottom flask equipped with a mechanical stirrer, thermometer, and dropping funnel under an atmosphere of nitrogen was added THF (25 mL), methylhydrazine (2.53 g, 2.92 mL, 55 mmol), and triethylamine (4.35 g, 5.99 mL, 55 mmol). The temperature of the mixture was lowered to approximately 5 °C and 4-methylbenzenesulfonyl chloride (9.5 g, 50 mmol) dissolved in 25 mL of THF was added dropwise at a rate such that the temperature did not rise over 10 °C. The cooling bath was removed and the mixture was allowed to stir overnight at RT. Ether (100 mL) was added to the reaction mixture and the resulting slurry was washed with water (50 mL), brine (50 mL), dried (Na<sub>2</sub>SO<sub>4</sub>), and the solvent was removed *in vacuo* to give 9.5 g (95 % yield) of the title product. <sup>1</sup>H NMR (CDCl<sub>3</sub>) δ 7.7 (d, 2H), 7.4 (d, 2H), 3.6 (s, b, 2H), 2.8 (s, 3H), 2.4 (s, 3H).

B. 1-(2-fluoro-6-chlorobenzoyl)-2-methyl-2-(4-methylphenyl)sulfonyl hydrazine

Into a 100 mL three necked round bottom flask equipped with a magnetic stirrer, thermometer, and condenser, under an atmosphere of nitrogen, was added 2-methyl-2-[(4-methylphenyl)sulfonyl] hydrazine (9.5 g, 47.4 mmol), pyridine (3.75 g, 3.83 mL, 47.4 mmol), 4-dimethylaminopyridine (0.30 g, 2.4 mmol) and acetonitrile (50 mL). The resulting mixture was cooled to 10 °C and 2-fluoro-6-chlorobenzoyl chloride (8.99 g, 45.2 mmol) dropwise at a rate such that the temperature did not rise above 10 °C. The cooling bath was removed and the

mixture was allowed to stir at RT overnight. The solvent was removed from the reaction mixture *in vacuo* to give an off white solid and the solid was diluted with methylene chloride and extracted with 1N HCl (20 mL), saturated sodium chloride solution (20 mL), dried ( $\text{Na}_2\text{SO}_4$ ), and the solvent removed *in vacuo* to give an off-white solid. This material was recrystallized from ethyl acetate to give 5.6 g (34.7% yield) of 1-(2-fluoro-6-chlorobenzoyl)-2-methyl-2-[(4-methylphenyl)sulfonyl] hydrazine as a crystalline solid: mp 135-136 °C;  $^1\text{H}$  NMR ( $\text{CDCl}_3$ )  $\delta$  7.6-7.8 (m, 3H), 7.0-7.4 (m, 5H), 3.4 and 3.0 (singlets combined for 3H), 2.4 and 2.45 (singlets combined for 3H);  $^{13}\text{C}$  ( $\text{CDCl}_3$ ) 160.2, 144.7, 133.6, 132.7, 132.6, 132.1, 131.9, 130.2, 129.6, 129.0, 128.7, 125.7, 114.6, 114.3, 37.7, 21.7. Anal. Calcd. For  $\text{C}_{15}\text{H}_{14}\text{ClFN}_2\text{O}_3\text{S}$ : C, 50.49; H, 3.95; N, 7.85. Found: C, 50.55; H, 3.89; N, 7.81.

C. N-(4-methylbenzenesulfonyl)-N-methyl-(2-fluoro-6-chlorobenz)hydrazonoyl chloride

Into a 250 mL one necked round bottom flask equipped with magnetic stirrer and condenser, under an atmosphere of nitrogen, was added 1-(2-fluoro-6-chlorobenzoyl)-2-methyl-2-[(4-methylphenyl)sulfonyl]hydrazine (5.0 g, 14.0 mmol), ethylene dichloride (50 mL), and phosphorus pentachloride (3.2 g, 21.6 mmol). The mixture was heated to reflux and was maintained at that temperature for approximately 30 min. The progress of the reaction was monitored by TLC using 50/50 ethyl acetate/hexanes (starting material  $R_f=0.55$ ; product  $R_f=0.70$ ). The solvent was removed *in vacuo* and the residue dissolved in methylene chloride (100 mL) and carefully diluted with water. The organic phase was washed with water, brine, dried ( $\text{Na}_2\text{SO}_4$ ) and the solvent removed *in vacuo* to give a yellow oil which solidified upon cooling. The solids were

slurried in hexane/ethyl acetate (70/30) and removed via filtration to give 4.5 g (87 % yield) of N-(4-methylbenzenesulfonyl)-N-methyl-(2-fluoro-6-chlorobenz)hydrazonoyl chloride. MP 98-99 °C ; <sup>1</sup>H NMR (CDCl<sub>3</sub>) δ 7.8 (d, 2H), 7.2-7.5 (m, 4H), 7.1 (m, H), 3.1 (s, 3H), 2.5 (s, 3H); <sup>13</sup>C NMR (CDCl<sub>3</sub>) 161.6, 158.5, 144.6, 140.7, 133.6, 132.1, 131.9, 131.1, 129.5, 125.6, 125.5, 114.7, 114.4, 38.1, 21.6. Anal. Calcd. for C<sub>15</sub>H<sub>13</sub>Cl<sub>2</sub>FN<sub>2</sub>O<sub>2</sub>S: C, 48.01; H, 3.49; N, 7.47. Found: C, 48.10; H, 3.49; N, 7.47.

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## Example 11

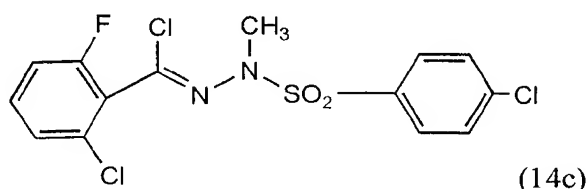
3-(2-chloro-6-fluorophenyl)-5-(3,4,5-trichlorothiophen-2-yl)-1-methyl[1,2,4]triazole (Compound 2)

Into a 50 mL necked flask equipped with a magnetic stirrer and thermometer, under an atmosphere of nitrogen, was added N-(4-methylbenzenesulfonyl)-N-methyl-(2-fluoro-6-chlorobenz)hydrazonoyl chloride (Example 10) (1.4 g, 3.73mmol), 2-cyano-3,4,5-trichlorothiophene (0.83 g, 3.92 mmol), aluminum chloride (0.53 g, 4.0 mmol), and o-dichlorobenzene (10 mL). The flask containing the reaction mixture was immersed into a preheated oil bath maintained at 120 °C. Within 8 minutes the temperature of the reaction mixture rose to 109 °C and heating was continued for an additional 30 min. The flask was removed from the oil bath. When the stirred reaction mixture reached approximately 80 °C and was poured into a solution (10 g of sodium hydroxide with 10 mL of ice) with rapid stirring and was then added to methylene chloride (50 mL) for extraction. The organic phase was washed with water (25 mL), brine, dried (Na<sub>2</sub>SO<sub>4</sub>), and the solvent removed in vacuo. The o-dichlorobenzene was removed from the residue via kugelrohr distillation to give a residue which was dissolved in ether (20 mL), stirred with activated carbon at RT, filtered, and the solvent was removed in vacuo

from the filtrate. The solids were recrystallized from ethyl acetate to give 0.9 g (60.8 % yield) of 3-(2-chloro-6-fluorophenyl)-5-(3,4,5-trichlorothien-2-yl)-1-methyl[1,2,4]triazole as a crystalline solid: >97% pure by  
5 HPLC analysis;  $^1\text{H}$  NMR ( $\text{CDCl}_3$ )  $\delta$  7.2-7.4 (m, 2H), 7.0-7.4 (m, 1H), 4.0 (s, 3H).

## Example 12

This Example shows preparation of the benzhydrazonoyl chloride of formula (14c):

A. 2-methyl-2-[(4-chlorophenyl)sulfonyl] hydrazine

Into a 100 mL three necked round bottom flask equipped with a mechanical stirrer, thermometer, and dropping funnel under an atmosphere of nitrogen was added THF (25 mL), methylhydrazine  
15 (2.53 g, 2.92 mL, 55 mmol), and triethylamine (4.35 g, 43 mmol). The temperature of the mixture was lowered to approximately 5 °C and crude 4-chlorophenylsulfonyl chloride (10.5 g, 50 mmol, contained 4-chlorophenyl sulfonic acid) dissolved in 25 mL of THF was added dropwise at a rate such  
20 that the temperature did not rise over 10 °C. The cooling bath was removed and the mixture was allowed to warm to stir overnight at RT. Ether (100 mL) was added to the reaction mixture and the resulting slurry was washed with water (50 mL), brine (50 mL), dried ( $\text{Na}_2\text{SO}_4$ ), and the solvent was removed  
25 *in vacuo* to give 5.4 g (49 % yield) of 2-methyl-2-[(4-chlorophenyl)sulfonyl] hydrazine.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ )  $\delta$  7.8 (d, 2H), 7.6 (d, 2H), 3.6 (s, b, 2H), 2.9 (s, 3H).

B. 1-(2-fluoro-6-chlorobenzoyl)-2-methyl-2-[(4-chlorophenyl)sulfonyl] hydrazine

Into a 100 mL three necked round bottom flask equipped with a magnetic stirrer and condenser, under an atmosphere of nitrogen, was added 2-methyl-2-[(4-chlorophenyl)sulfonyl] hydrazine (5.4 g, 24.5 mmol), pyridine (1.93 g, 1.98 mL, 24.5 mmol), 4-dimethylaminopyridine (0.15 g, 1.2 mmol), and acetonitrile (50 mL). The mixture was cooled to 10 °C and 2-fluoro-6-chlorobenzoyl chloride (4.65 g, 23.4 mmol) was added dropwise at a rate such that the temperature did not rise above 10 °C. The mixture was allowed to stir at RT overnight. The solvent was removed from the reaction mixture *in vacuo* to give an off white solid and the solid was diluted with methylene chloride and extracted with 1N HCl (20 mL), saturated sodium bicarbonate (20 mL), dried (Na<sub>2</sub>SO<sub>4</sub>), and the solvent removed *in vacuo* to give 1-(2-fluoro-6-chlorobenzoyl)-2-methyl-2-[(4-chlorophenyl)sulfonyl] hydrazine as a yellow oil which was used without purification.

C. N-(4-chlorophenylsulfonyl)-N-methyl-(2-fluoro-6-chlorobenz)hydrazonoyl chloride

Into a 250 mL one necked round bottom flask equipped with magnetic stirrer and condenser, under an atmosphere of nitrogen, was added 1-(2-fluoro-6-chlorobenzoyl)-2-methyl-2-(4-chlorophenylsulfonyl) hydrazine (7.3 g, 16.8 mmol, 87% pure by HPLC), ethylene dichloride (50 mL), and phosphorus pentachloride (6.03 g, 28.9 mmol). The mixture was heated to reflux and was maintained at that temperature for approximate 30 min. The solvent was removed *in vacuo* and the residue dissolved in methylene chloride (100 mL) and carefully diluted with water. The organic phase was washed with water, brine, dried (Na<sub>2</sub>SO<sub>4</sub>) and the solvent removed *in vacuo* to give a

yellow oil which was dissolved in hot ethyl acetate to which was added hexanes to the cloud point and upon cooling with scratching solids were formed. The solids were recrystallized from hexane/ethyl acetate (70/30) and seeded to give 2.7 g (40.9 % yield) of N-(4-chlorophenylsulfonyl)-N-methyl-(2-fluoro-6-chlorobenz)hydrazonoyl chloride. MP 100-101 °C; <sup>1</sup>H NMR (CDCl<sub>3</sub>) δ 7.8 (d, 2H), 7.2-7.5 (m, 4H), 7.1 (m, H), 3.1 (s, 3H), 2.5 (s, 3H). Anal. Calcd. for C<sub>14</sub>H<sub>10</sub>Cl<sub>3</sub>FN<sub>2</sub>O<sub>2</sub>S: C, 42.50; H, 2.55; N, 7.08. Found: C, 42.56; H, 2.47; N, 6.99.

### Example 13

#### 3-(2-chloro-6-fluorophenyl)-5-(3,4,5-trichlorothiophen-2-yl)-1-methyl[1,2,4]triazole (Compound 2)

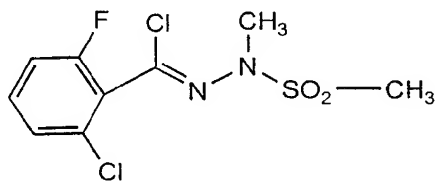
Into a 50 mL necked flask equipped with a magnetic stirrer and thermometer, under an atmosphere of nitrogen, was added N-(4-chlorophenylsulfonyl)-N-methyl-(2-fluoro-6-chlorobenz)-hydrazonoyl chloride (Preparation 4) (2.7 g, 6.8 mmol), 2-cyano-3,4,5-trichlorothiophene (1.44 g, 6.8 mmol), aluminum chloride (1.18 g, 8.9 mmol), and o-dichlorobenzene (20 mL). The flask containing the reaction mixture was immersed into a preheated oil bath maintained at 120 °C. Within 8 minutes the temperature of the reaction mixture rose to 109 °C and heating was continued for an additional 50 min. The flask was removed from the oil bath. When the stirred reaction mixture reached approximately 80 °C and was poured into a solution 20 g of sodium hydroxide with 20 mL of ice) with rapid stirring and was then added to methylene chloride (100 mL) for extraction. The organic phase was washed with water, brine, dried (Na<sub>2</sub>SO<sub>4</sub>), and the solvent removed *in vacuo*. The o-dichlorobenzene was removed from the residue via kugelrohr distillation to give a residue which was dissolved in ether (50 mL), stirred with activated carbon at RT, filtered, and the solvent was removed *in vacuo* from the

filtrate. The resulting off white solids were recrystallized to give 1.8 g (26.6 % yield) of 3-(2-chloro-6-fluorophenyl)-5-(3,4,5-trichlorothien-2-yl)-1-methyl[1,2,4]triazole as a crystalline solid: >98% pure by HPLC analysis.

5

## Example 14

This Example shows preparation of the benzhydrazonoyl chloride of formula (14d):



(14d)

10 A. 2-Methyl-2-methanesulfonyl hydrazine

Into a 250 mL three necked round bottom flask equipped with a mechanical stirrer, thermometer, and dropping funnel under an atmosphere of nitrogen was added THF (100 mL), methyl hydrazine (5.07 g, 5.85 mL, 110 mmol), and triethylamine (11.1 g, 15.3 mL, 110 mmol). The temperature of the mixture was lowered to approximately 5 °C and methanesulfonyl chloride (11.4 g, 7.73 mL, 100 mmol) was added dropwise at a rate such that the temperature did not rise above 10°C. The cooling bath was removed and the mixture was allowed to warm to RT and was stirred over night. Ether (100 mL) was added to the reaction mixture and washed with water (50 mL), brine (50 mL), dried ( $Na_2SO_4$ ), and the solvent was removed *in vacuo* to give 2.4 g (19.3 % yield) of 2-methyl-2-methanesulfonyl hydrazine.  $^1H$  NMR ( $CDCl_3$ )  $\delta$  3.8 (s, b, 2H), 2.95 (s, 3H), 2.9 (s, 3H).

25

B. 1-(2-fluoro-6-chlorobenzoyl)-2-methyl-2-methanesulfonyl hydrazine



Into a 50-mL was added 2-methyl-2-methanesulfonyl hydrazine (2.4 g, 19.3 mmol), pyridine (1.5 g, 1.55 mL, 19.3 mmol), and acetonitrile (20 mL), under an atmosphere of nitrogen. The mixture was cooled to approximately 0 °C and 2-chloro-6-fluorobenzoyl chloride (3.77 g, 19.3 mmol) was added dropwise. The mixture was allowed to warm to RT and to stir for approximately 3 hr with monitoring by TLC using 50/50 ethyl acetate/hexanes (Rf=0.7 for 2-chloro-6-fluorobenzoyl chloride and Rf=0.44 for product). The solvent was removed from the reaction mixture in vacuo, the residue was diluted with methylene chloride, and extracted with 1N hydrochloric acid (10 mL), saturated sodium bicarbonate, dried (MgSO<sub>4</sub>), and the solvent removed in vacuo to give 4.6 g of a semi-solid. This material was recrystallized from ethyl acetate to give 1.8 g (33.3% yield) of 1-(2-fluoro-6-chlorobenzoyl)-2-methyl-2-methanesulfonyl hydrazine as a crystalline solid. MP 135-136 °C ; <sup>1</sup>H NMR (CDCl<sub>3</sub>) δ 7.8 (s, b, 1H), 7.2-7.4 (m, 3H), 3.4 (s, 3H), 3.15 (s, 3H). Anal. Calcd. for C<sub>9</sub>H<sub>10</sub>ClFN<sub>2</sub>O<sub>3</sub>S: C, 38.51; H, 3.59; N, 9.98. Found: C, 38.28, H, 3.41; N, 9.48.

C. N-methanesulfonyl-N-methyl-(2-fluoro-6-chlorobenz)hydrazonoyl chloride

Into a 250 mL one necked round bottom flask equipped with magnetic stirrer and condenser, under an atmosphere of nitrogen, was added 1-(2-fluoro-6-chlorobenzoyl)-2-methyl-2-methanesulfonyl hydrazine (1.6 g, 7.12 mmol), ethylene dichloride (25 mL), and phosphorus pentachloride (2.2 g, 10.7 mmol). The mixture was heated to reflux and was maintained at that temperature for approximate 30 min. The progress of the reaction was monitored by TLC using 50/50 ethyl acetate/hexanes (starting material Rf=0.44; product Rf=0.60. The solvent was removed in vacuo and the residue dissolved in

methylene chloride (50 mL) and carefully diluted with water. The organic phase was washed with water (2 X 50 mL), brine (50 mL), dried ( $\text{Na}_2\text{SO}_4$ ) and the solvent removed in vacuo to give a colorless oil which solidified upon cooling. The solids were  
5 recrystallized from hexanes/ethyl acetate (70/30) to give 1.2 g (56.6 % yield) of N-methanesulfonyl-N-methyl-(2-fluoro-6-chlorobenz)hydrazonoyl chloride. MP 83-85 °C ; 93% pure by  
HPLC analysis:  $^1\text{H}$  NMR ( $\text{CDCl}_3$ )  $\delta$  7.8 (d, 2H), 7.0-7.4 (m, 3H), 3.45 and 3.35 (singlets adding to 3H), 3.15 and 3.05 (singlets  
10 adding to 3H).

## Example 15

3-(2-chloro-6-fluorophenyl)-5-(3,4,5-trichlorothiophen-2-yl)-1-methyl[1,2,4]triazole (Compound 2)

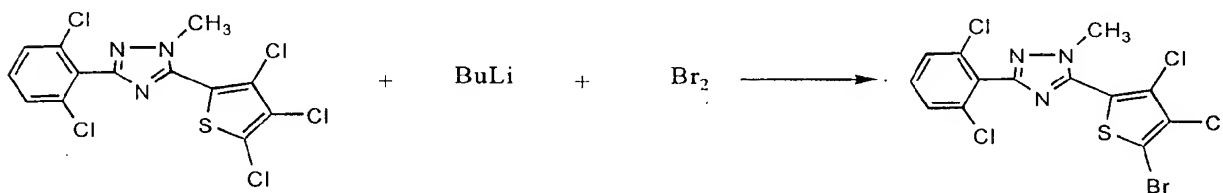
15 Into a 50 mL necked flask equipped with a magnetic stirrer and thermometer, under an atmosphere of nitrogen, was added N-methanesulfonyl-N-methyl-(2-fluoro-6-chlorobenz)hydrazonoyl chloride (Example 14) (1.0 g, 3.3 mmol),  
2-cyano-3,4,5-trichlorothiophene (0.78 g, 3.7 mmol), aluminum  
20 chloride (0.49 g, 3.7 mmol), and o-dichlorobenzene (10 mL). The flask containing the reaction mixture was immersed into a preheated oil bath maintained at 120 °C. Within 10 minutes the temperature of the reaction mixture rose to 111 °C and heating was continued for an additional 90 min. GC analysis  
25 indicated that the conversion of the reaction was approximately 66%. The flask was removed from the oil bath. When the stirred reaction mixture reached approximately 90 °C and was poured into a solution (10 g of sodium hydroxide with 10 mL of ice) with rapid stirring and was then added to  
30 methylene chloride (50 mL) for extraction. The organic phase was washed with water (25 mL), brine, dried ( $\text{Na}_2\text{SO}_4$ ), and the solvent removed in vacuo. The o-dichlorobenzene was removed

from the residue via kugelrohr distillation to give a residue which was dissolved in ether (20 mL), stirred with activated carbon at RT, filtered, and the solvent was removed *in vacuo* from the filtrate to give 0.4 g of 3-(2-chloro-6-fluorophenyl)-5-(3,4,5-trichlorothien-2-yl)-1-methyl[1,2,4]triazole contaminated with 3,4,5-trichloro-2-cyanothiophene.

Compounds of the invention can be modified using conventional processes to provide other compounds of the invention, as illustrated in Examples 16 and 17.

#### Example 16

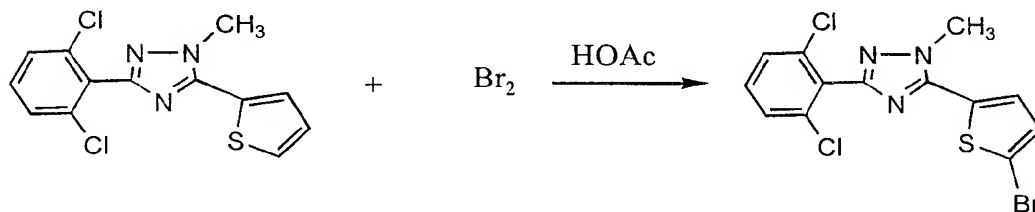
3-(2,6-dichlorophenyl)-5-(5-bromo-3,4-dichlorothien-2-yl)-1-methyl[1,2,4]triazole (Compound 46)



n-Butyllithium (2.5 mmol) was slowly added to a solution of 3-(2,6-dichlorophenyl)-5-(3,4,5-trichlorothien-2-yl)-1-methyl[1,2,4]triazole (325 mg, 1.0 mmol) in THF at  $-78^{\circ}\text{C}$ , and the mixture was stirred one hour. Bromine (3.0 mmol) was added and the reaction stirred two hours, then diluted with water, extracted with ether, washed with brine, and dried over magnesium sulfate. The product was concentrated to 620 mg of dark oil. This was chromatographed ( $\text{SiO}_2$ , 10% EtOAc-Hex) to give 79 mg of the title product as a pink solid. MP  $197-199^{\circ}\text{C}$ .

## Example 17

3-(2,6-dichlorophenyl)-5-(5-bromothiophen-2-yl)-1-methyl  
[1,2,4]triazole (Compound 47)

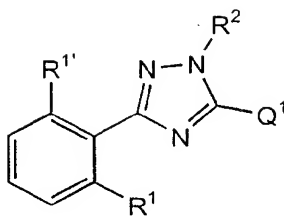


5        3-(2,6-dichlorophenyl)-5-(thien-2-yl)-1-methyl-  
 [1,2,4]triazole (75 mg, 0.242 mmol) and bromine (39 mg, 0.242  
 mmol) were combined in glacial acetic acid (3mL) and the  
 mixture was stirred overnight at room temperature, then heated  
 to 95°C for five hours. The reaction mixture was poured into  
 10 a saturated sodium bicarbonate solution, extracted with ether,  
 and dried over magnesium sulfate. Chromatography (SiO<sub>2</sub>, 25%  
 Hex-CH<sub>2</sub>Cl<sub>2</sub>) delivered the title product as a waxy white solid.  
 MP 130-132°C.

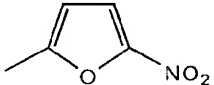
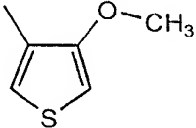
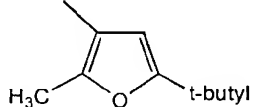
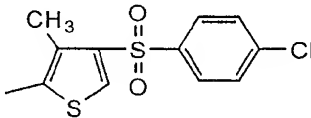
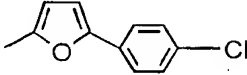
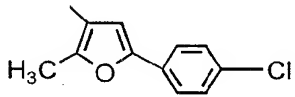
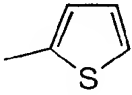

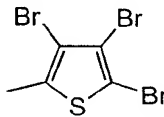
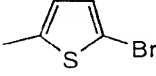
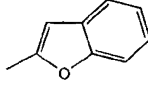
15        Phytologically acceptable acid addition salts of the  
 compounds of formula (1) are also within the scope of the  
 invention. For example, boron tetrafluoride, hydrogen  
 chloride, hydrogen bromide, hydrogen iodide, hydrogen sulfate,  
 or organic acid salts may be used.

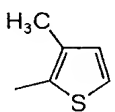
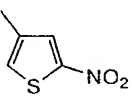
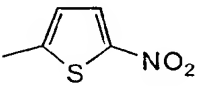
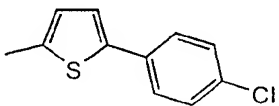
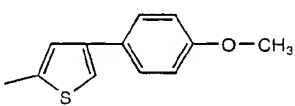
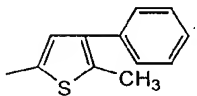
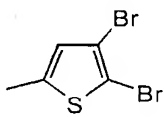
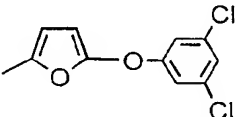
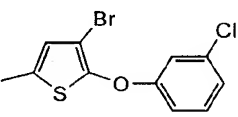
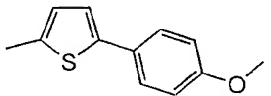
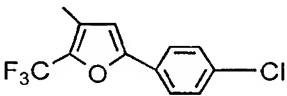
20        The compounds identified in the following Table 1 were  
 prepared and tested.

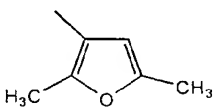
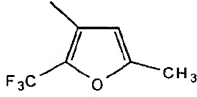
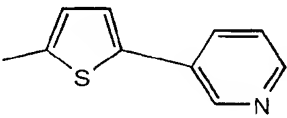
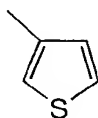
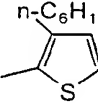
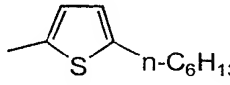
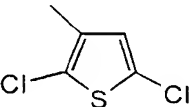
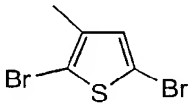
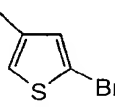
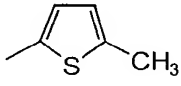
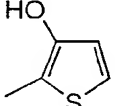
Table 1.



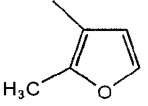
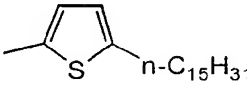
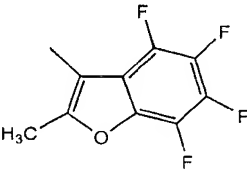
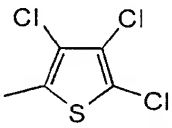
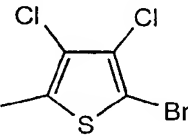
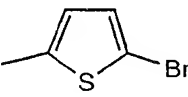
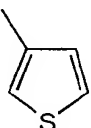
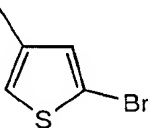
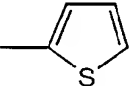
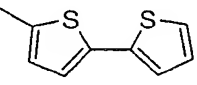
compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
1	Cl	F	CH <sub>3</sub>		112-113	A	C	E
2	Cl	F	CH <sub>3</sub>		127-128	A	A	B
3	Cl	F	CH <sub>3</sub>		145-147	G	A	
4	F	F	CH <sub>3</sub>		oil	F		
5	F	F	CH <sub>3</sub>		137-138	A	F	B
6	F	F	CH <sub>3</sub>		104-106	C		E
7	F	F	CH <sub>3</sub>		60-61	A	A	C
8	F	F	CH <sub>3</sub>		176-177	F		

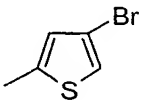
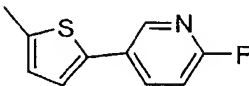
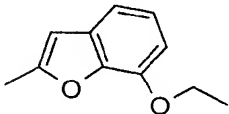
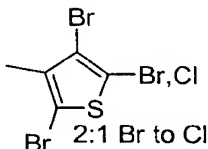
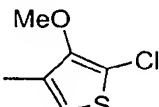
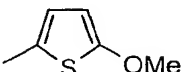
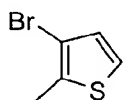
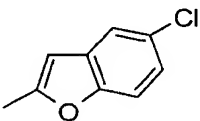
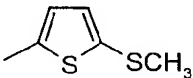
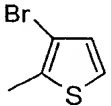
compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
9	F	F	CH <sub>3</sub>		172-173	F		
10	F	F	CH <sub>3</sub>		oil	A	A	C
11	F	F	CH <sub>3</sub>		76-78	F		
12	F	F	CH <sub>3</sub>		183-185	G	A	G
13	F	F	CH <sub>3</sub>		171-173	B	A	
14	F	F	CH <sub>3</sub>		146-148	D		
15	Cl	F	CH <sub>3</sub>		109-111	A	F	D
16	F	F	CH <sub>3</sub>		101-102	A	A	G
17	F	F	CH <sub>3</sub>		153-154	A	A	F
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19	F	F	CH <sub>3</sub>		98-100	A		

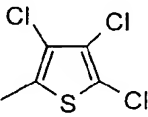
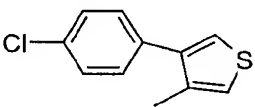
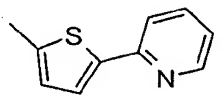
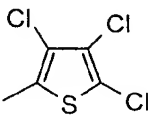
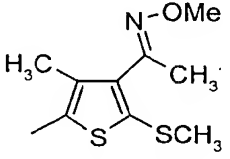
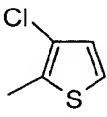
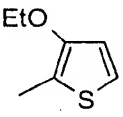
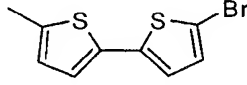
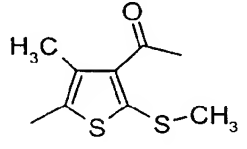
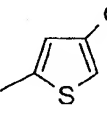
cmpd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
20	Cl	F	CH <sub>3</sub>		80-82	A	D	A
21	Cl	F	CH <sub>3</sub>		113-115	C	A	G
22	Cl	F	CH <sub>3</sub>		129-130	G	A	F
23	F	F	CH <sub>3</sub>		127-129	F	G	D
24	F	F	CH <sub>3</sub>		106-108	F	B	F
25	F	F	CH <sub>3</sub>		135-136	F	A	G
26	F	F	CH <sub>3</sub>		136-138	A	C	F
27	F	F	CH <sub>3</sub>		oil	E		
28	F	F	CH <sub>3</sub>		142-144	F	A	F
29	F	F	CH <sub>3</sub>		100-101	A	B	F
30	F	F	CH <sub>3</sub>		159-161	G		

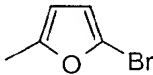
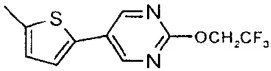
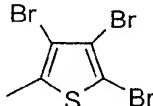
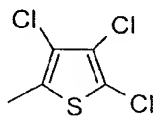
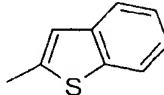
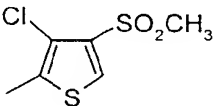
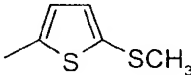
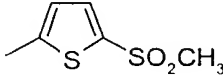
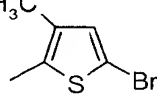
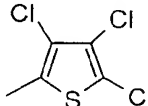
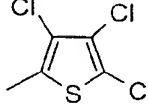
compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
31	F	F	CH <sub>3</sub>		oil	C		
32	F	F	CH <sub>3</sub>		53-55	A		
33	F	F	CH <sub>3</sub>		169-170	F		
34	Cl	F	CH <sub>3</sub>		100-101	A	G	
35	F	F	CH <sub>3</sub>		oil	D		
36	F	F	CH <sub>3</sub>		oil	F		
37	Cl	F	CH <sub>3</sub>		oil	A	E	
38	Cl	F	CH <sub>3</sub>		138-139	A		
39	Cl	F	CH <sub>3</sub>		83-84	A	B	
40	F	F	CH <sub>3</sub>		97-99	D		
41	F	F	CH <sub>3</sub>		146-147	G		

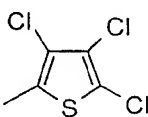
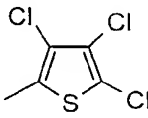
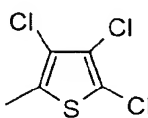
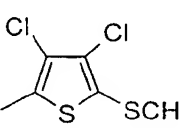
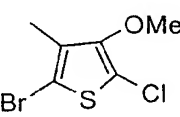
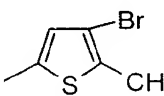
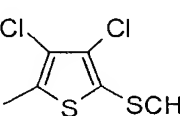
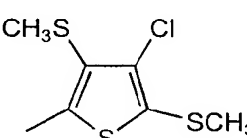
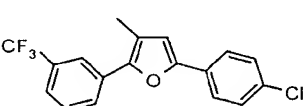
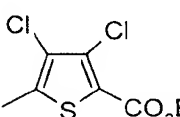


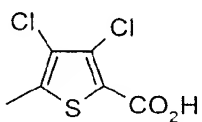
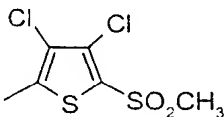
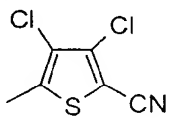
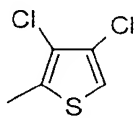
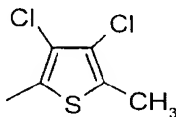
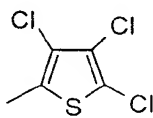
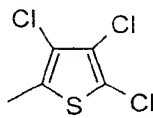
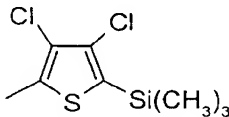
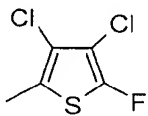
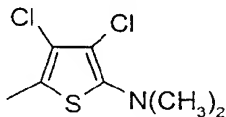
compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
42	F	F	CH <sub>3</sub>		oil	A		
43	F	F	CH <sub>3</sub>		63-64	F		
44	F	F	CH <sub>3</sub>		115-117	F		
45	Cl	Cl	CH <sub>3</sub>		151-152	A		
46	Cl	Cl	CH <sub>3</sub>		197-199	A		
47	Cl	Cl	CH <sub>3</sub>		130-132	A		
48	Cl	Cl	CH <sub>3</sub>		oil	A		
49	Cl	Cl	CH <sub>3</sub>		yellow oil	A		
50	Cl	Cl	CH <sub>3</sub>		161-163	A		
51	F	F	CH <sub>3</sub>		116-118	G		

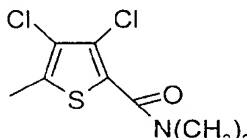
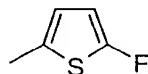
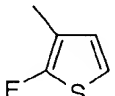
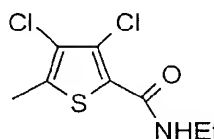
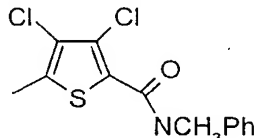
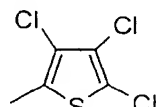
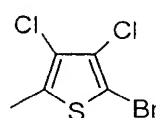
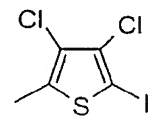
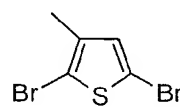
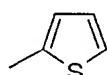
compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
52	F	F	CH <sub>3</sub>		153-155	D		
53	F	F	CH <sub>3</sub>		147-148	G		
54	F	F	CH <sub>3</sub>		88-90	F		
55	F	F	CH <sub>3</sub>	 Br 2:1 Br to Cl	oil	G		
56	F	F	CH <sub>3</sub>		83-85	B		
57	F	F	CH <sub>3</sub>		125-126	A	G	
58	Cl	F	CH <sub>3</sub>		99-100	B		
59	F	F	CH <sub>3</sub>		150-152	D		
60	Cl	F	CH <sub>3</sub>		oil	B		
61	F	F	CH <sub>3</sub>		98-99	A		

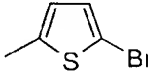
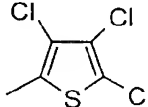
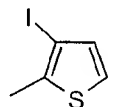
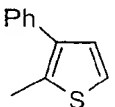
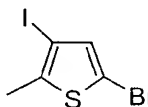
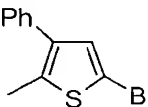
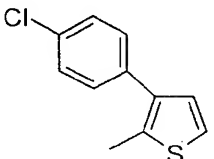
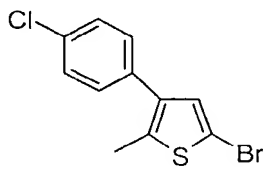
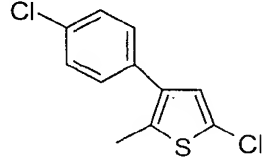
compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
62	Cl	F	CH <sub>2</sub> Cl		105-106	A		
63	F	F	CH <sub>3</sub>		oil	G		
64	F	F	CH <sub>3</sub>		144-146	G		
65	Cl	F	CHCl <sub>2</sub>		114-115	F		
66	F	F	CH <sub>3</sub>		oil	A		
67	Cl	F	CH <sub>3</sub>		87-88	A		
68	Cl	F	CH <sub>3</sub>		111-113	G		
69	F	F	CH <sub>3</sub>		127-129	F		
70	F	F	CH <sub>3</sub>		oil	F		
71	Cl	F	CH <sub>3</sub>		130-132	A		

compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WP <sup>*</sup>
72	F	F	CH <sub>3</sub>		oil	F		
73	F	F	CH <sub>3</sub>		177	F		
74	Cl	F	CH <sub>3</sub>		206-208	A		
75	Cl	F	CH <sub>2</sub> C H <sub>3</sub>		83-84	A		
76	F	F	CH <sub>3</sub>		145-147	C		
77	F	F	CH <sub>3</sub>		141-142	F		
78	F	F	CH <sub>3</sub>		oil	B		
79	F	F	CH <sub>3</sub>		157-158	F		
80	Cl	F	CH <sub>3</sub>		117-118	A		
81	Cl	F	CH <sub>2</sub> CH <sub>2</sub> CH <sub>3</sub>		116-117	D		
82	Cl	F	CH <sub>2</sub> C N		oil	F		

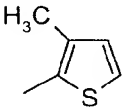
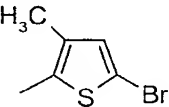
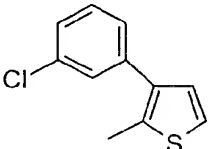
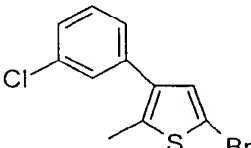
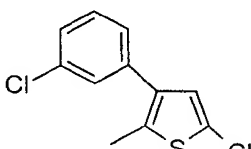
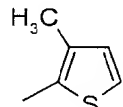
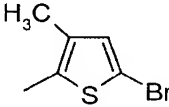
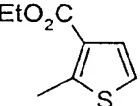
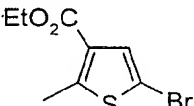
compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
83	Cl	F	CHClF		oil	D		
84	Cl	F	CH <sub>2</sub> SC H <sub>3</sub>		oil	F		
85	Cl	F	CH <sub>2</sub> O CH <sub>3</sub>		oil	A		
86	Cl	F	CH <sub>2</sub> SC H <sub>3</sub>		oil	A		
87	F	F	CH <sub>3</sub>		oil	E		
88	F	F	CH <sub>3</sub>		122	F		
89	Cl	F	CH <sub>3</sub>		101.5- 102	E		
90	Cl	F	CH <sub>3</sub>		115- 116.5	A		
91	F	F	CH <sub>3</sub>		157-159	D		
92	Cl	F	CH <sub>3</sub>		109.5- 112	B		

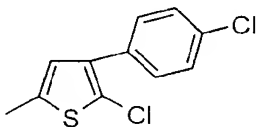
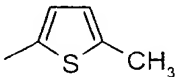
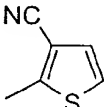
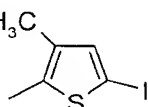
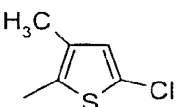
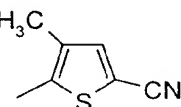
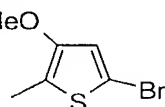
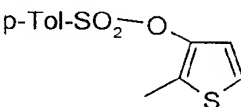
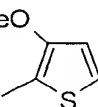
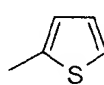
compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
93	Cl	F	CH <sub>3</sub>		267-270 dec	F		
94	Cl	F	CH <sub>3</sub>		178-180	F		
95	Cl	F	CH <sub>3</sub>		95-97	B		
96	Cl	F	CH <sub>3</sub>		118-120	A		
97	Cl	F	CH <sub>3</sub>		82-83.5	B		
98	Cl	F	CH <sub>2</sub> SO <sub>2</sub> CH <sub>3</sub>		207-209	F		
99	Cl	F	CH <sub>2</sub> F		oil	A		
100	Cl	F	CH <sub>3</sub>		133-135	B		
101	Cl	F	CH <sub>3</sub>		128-131	A		
102	Cl	F	CH <sub>3</sub>		126-131	B		

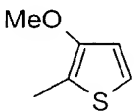
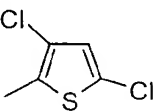
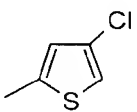
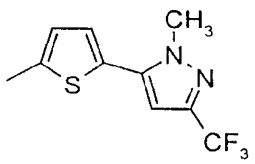
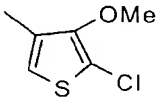
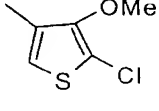
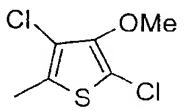
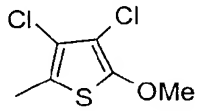
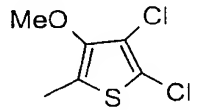
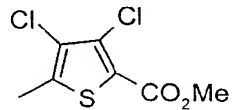
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103	Cl	F	CH <sub>3</sub>		131-133	C		
104	Cl	F	CH <sub>3</sub>		113-115	A		
105	Cl	F	CH <sub>3</sub>		59-64	A		
106	Cl	F	CH <sub>3</sub>		162.5-164	F		
107	Cl	F	CH <sub>3</sub>		oil	F		
108	H	Cl	CH <sub>3</sub>		162-164	C		
109	Cl	F	CH <sub>3</sub>		178-180	A		
110	Cl	F	CH <sub>3</sub>		218-223	A		
111	Cl	Cl	CH <sub>3</sub>		oil	C		
112	H	Cl	CH <sub>3</sub>		oil	B		

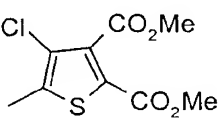
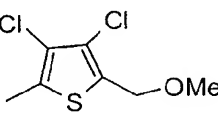
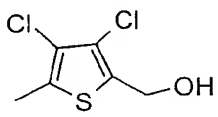
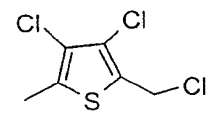
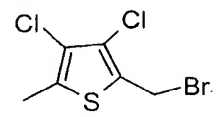
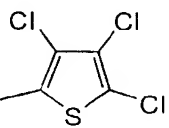
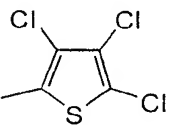
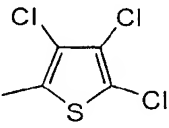
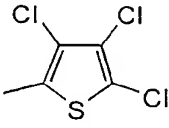
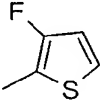
compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
113	H	Cl	CH <sub>3</sub>		oil	A		
114	H	CF <sub>3</sub>	CH <sub>3</sub>		155-156	F		
115	Cl	F	CH <sub>3</sub>		68-72	A		
116	Cl	F	CH <sub>3</sub>		107-110	F		
117	Cl	F	CH <sub>3</sub>		131-133	B		
118	Cl	F	CH <sub>3</sub>		oil	F		
119	Cl	F	CH <sub>3</sub>		oil	F		
120	Cl	F	CH <sub>3</sub>		oil	F		
121	Cl	F	CH <sub>3</sub>		oil	F		

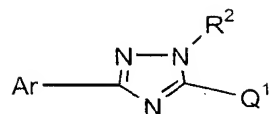


compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
122	F	F	CH <sub>3</sub>		97-99	B		
123	F	F	CH <sub>3</sub>		89-91	C		
124	Cl	F	CH <sub>3</sub>		oil	G		
125	Cl	F	CH <sub>3</sub>		oil	F		
126	Cl	F	CH <sub>3</sub>		oil	F		
127	Cl	Cl	CH <sub>3</sub>			C		
128	Cl	Cl	CH <sub>3</sub>			G		
129	Cl	F	CH <sub>3</sub>		oil	F		
130	Cl	F	CH <sub>3</sub>		103-106	F		

compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
131	Cl	F	CH <sub>3</sub>		157-158	F		
132	Cl	F	CH <sub>3</sub>		97-98	A		
134	Cl	F	CH <sub>3</sub>		113-115	G		
135	Cl	Cl	CH <sub>3</sub>			C		
136	Cl	Cl	CH <sub>3</sub>			B		
137	Cl	Cl	CH <sub>3</sub>			C		
138	Cl	F	CH <sub>3</sub>		121-122	F		
139	Cl	F	CH <sub>3</sub>		129-130	G		
140	Cl	F	CH <sub>3</sub>		oil	C		
141	OMe	Cl	CH <sub>3</sub>		oil	A		

compd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
142	OMe	Cl	CH <sub>3</sub>		146-148	F		
143	Cl	F	CH <sub>3</sub>		97-103	A		
144	F	F	CH <sub>3</sub>		92-94	A		
145	F	F	CH <sub>3</sub>		133-134	F		
146	Cl	F	CH <sub>3</sub>					
147	Cl	Cl	CH <sub>3</sub>					
148	Cl	F	CH <sub>3</sub>					
149	Cl	F	CH <sub>3</sub>					
150	Cl	F	CH <sub>3</sub>					
151	Cl	F	CH <sub>3</sub>					

cmpd no.	R <sup>1</sup>	R <sup>1'</sup>	R <sup>2</sup>	Q <sup>1</sup>	mp °C	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
152	Cl	F	CH <sub>3</sub>					
153	Cl	F	CH <sub>3</sub>					
154	Cl	F	CH <sub>3</sub>					
155	Cl	F	CH <sub>3</sub>					
156	Cl	F	CH <sub>3</sub>					
157	OCH <sub>3</sub>	Cl	CH <sub>3</sub>					
158	OCH <sub>3</sub>	F	CH <sub>3</sub>					
159	CH <sub>3</sub>	Cl	CH <sub>3</sub>					
160	CH <sub>3</sub>	F	CH <sub>3</sub>					
161	Cl	F	CH <sub>3</sub>					



compound number	Ar	R <sup>2</sup>	Q <sup>1</sup>	mp	CA <sup>†</sup>	TSSM <sup>‡</sup>	WF <sup>*</sup>
162		CH <sub>3</sub>		89-94	G		
163		CH <sub>3</sub>		187-191			
164		CH <sub>3</sub>		188-189	F		
165		CH <sub>3</sub>					
166		CH <sub>3</sub>					
167		CH <sub>3</sub>					
168		CH <sub>3</sub>					

CA<sup>†</sup> refers to activity at 50 ppm against cotton aphid,  
TSSM<sup>†</sup> refers to activity at 100 ppm against to-spotted spider  
mite, and

WF\* refers to activity at 800 ppm against whitefly. The test  
procedures are described below.

In each case the rating scale is as follows

% Control	Rating
91-100	A
81-90	B
71-80	C
61-70	D
51-60	E
less than 51	F
inactive	G

#### Insecticide and Miticide Utility

The compounds of the invention are also useful for the  
control of insects, mites, and aphids. Therefore, the present  
invention also is directed to a method for inhibiting an  
insect, mite, or aphid which comprises applying to a locus of  
the insect or mite an insect- or mite-inhibiting amount of a  
compound of formula (1).

The compounds are useful for reducing populations of  
insects and mites and are useful in a method of inhibiting an  
insect or mite population which comprises applying to a locus  
of the insect or mite an effective insect- or mite-  
inactivating amount of a compound of formula (1). The "locus"  
of insects or mites is a term used herein to refer to the  
environment in which the insects or mites live or where their  
eggs are present, including the air surrounding them, the food  
they eat, or objects which they contact. For example, plant-  
ingesting insects or mites can be controlled by applying the  
active compound to plant parts that the insects or mites eat,

particularly the foliage. It is contemplated that the compounds might also be useful to protect textiles, paper, stored grain, or seeds by applying an active compound to such substance. The term "inhibiting an insect or mite" refers to  
5 a decrease in the numbers of living insects or mites, or a decrease in the number of viable insect or mite eggs. The extent of reduction accomplished by a compound depends, of course, upon the application rate of the compound, the particular compound used, and the target insect or mite  
10 species. At least an inactivating amount should be used. The terms "insect-inactivating amount" and "mite-inactivating amount" are used to describe the amount, which is sufficient to cause a measurable reduction in the treated insect or mite, population. Generally an amount in the range from about 1 to  
15 about 1000 ppm active compound is used.

In a preferred embodiment, the present invention is directed to a method for inhibiting a mite or aphid which comprises applying to a plant an effective mite- or aphid-inactivating amount of a compound of formula (1).

20 Insecticidal test for cotton aphid (*Aphis gossypii*):

To prepare spray solutions, 2 mg of each test compound was dissolved into 2 mL of a 90:10 acetone:ethanol solvent. This 2 mL of chemical solution was added to 38 mL of water containing 0.05% Tween 20 surfactant to produce a 50 ppm spray  
25 solution.

Squash cotyledons were infested with cotton aphid (all life stages) 16-20 hours prior to application of spray solution. The solution was sprayed on both sides of each infested squash cotyledon (0.5 mL X 2 each side) with a  
30 sweeping action using a total of 2 mL of spray solution. The plants were allowed to air dry and held for 3 days in a controlled room at 26° C and 40 % RH after which time the test was graded. Grading was by actual count using a dissecting

microscope and comparison of test counts to the untreated check. Results are given in Table 1 as percent control based on population reduction versus the untreated.

Insecticidal test for two-spotted spider mite (*Tetranychus urticae*)

Ovicide Method:

Ten adult female two-spotted spider mites were placed on eight 2.2 cm leaf discs of cotton leaf, allowed to oviposit over 24 hours, and thereafter removed. The leaf discs were immersed in 100 ppm test solutions for 3 seconds, then allowed to dry with sixteen discs left untreated as a negative control. Discs were placed on an agar substrate and held at 24 °C and 90% relative humidity for 6 days. Percent control based on the number of hatched larvae on treated discs and the number on untreated discs is reported in Table 1.

Evaluation of Tests Compounds on Sweetpotato Whitefly (*Bemisia tabacia*) Under Laboratory Conditions

Sixteen mg of each test compound was dissolved by adding 5 ml of a 90:10 acetone:ethanol solvent mixture to the vial containing the sample compound. This solution was added to 15 ml of water containing 0.05% Tween 20 surfactant to produce 20 ml of an 800 ppm spray solution.

Five-week-old cotton plants reared in a greenhouse were stripped of all foliage except for the two uppermost true leaves that were greater than 5 cm in diameter. These plants were then placed into a laboratory colony of whiteflies for three days for oviposition by the colony females. All whiteflies were then removed from the test plants with pressurized air. The spray solution was then applied to the test plants with a hand-held syringe fitted with hollow cone nozzle. One mL spray solution was applied to each leaf top



and bottom for a total of 4 mL per plant. Four replications of each test compound utilized a total of 16 mL spray solution. Plants were air dried and then placed in a holding chamber (26° C and 40% RH) for 21 days. Compound efficacy was evaluated by counting, under a dissecting microscope, the number of spent pupal cases per leaf. A spent pupal case represents a whitefly egg that has undergone full development to achieve adult status, thus indicating lack of control.

Percent control based on reduction of spent pupal cases of a test compound compared to solution-only (no test compound) sprayed plants is reported in Table 1.

#### Seed Treatment Test vs. Cotton Aphid on Squash

Four-inch pots were filled two-thirds full with sieved soil mix (70 sand/30 pulverized soil). Compound was applied as a suspension to the center in 4 ml volume. Squash seed placed in the applied chemical zone was covered with soil mix 1 cm deep. The pots were irrigated and maintained in the greenhouse for ten days, at which time each plant had an expanded full leaf available for infestation. The plants were then infested with aphids. Four days later, aphid counts were made. Results are summarized in the following tables:

#### **Cotton aphid counts on Squash - cotyledon**

Compound	Rate mg/pot	Mean	SD	% Ctrl.
Compound 2	12.8	7.0	6.2	93.3
	3.2	7.3	4.0	93.1
	0.8	48.0	42.1	54.4
Compound 45	12.8	80.0	37.7	24.0
	3.2	50.3	14.8	52.3
	0.8	165.5	8.6	0
Imidacloprid (control)	0.8	0	0	100
	0.2	0	0	100
	0.05	2.5	3.5	97.6
Check	0	105.3	45.8	0

Cotton aphid counts on Squash - 1<sup>st</sup> True Leaf

Compd	Rate mg/pot	Mean	SD	% Ctrl.
Compound 2	12.8	3.3	2.6	96.1
	3.2	3.5	2.4	95.9
	0.8	12.0	14.2	85.8
Compound 45	12.8	8.3	5.1	90.2
	3.2	18.8	12.5	77.8
	0.8	40.8	18.9	51.7
Imidacloprid (control)	0.8	0.0	0.0	100.0
	0.2	9.8	6.5	88.4
	0.05	35.0	21.2	58.5
Check	0	84.4	53.2	0

In addition to being effective against mites, aphids, and insects when applied to foliage, compounds of formula (1) have exhibited systemic activity. Accordingly, another aspect of the invention is a method of protecting a plant from insects which comprises treating plant seed prior to planting it, treating soil where plant seed is to be planted, or treating soil at the roots of a plant after it is planted, with an effective amount of a compound of formula (1).

10

Compositions

The compounds of this invention are applied in the form of compositions which are important embodiments of the invention, and which comprise a compound of this invention and a phytologically-acceptable inert carrier. The compositions are either concentrated formulations which are dispersed in water for application, or are dust or granular formulations which are applied without further treatment.

The compositions are prepared according to procedures and formulae which are conventional in the agricultural chemical art, but which are novel and important because of the presence therein of the compounds of this invention. Some description of the formulation of the compositions will be given, however, to assure that agricultural chemists can readily prepare any desired composition.

The dispersions in which the compounds are applied are most often aqueous suspensions or emulsions prepared from concentrated formulations of the compounds. Such water-soluble, water-suspendable or emulsifiable formulations are either solids, usually known as wettable powders, or liquids usually known as emulsifiable concentrates or aqueous suspensions. Wettable powders, which may be compacted to form water dispersible granules, comprise an intimate mixture of the active compound, an inert carrier, and surfactants. The concentration of the active compound is usually from about 10% to about 90% by weight. The inert carrier is usually chosen from among the attapulgite clays, the montmorillonite clays, the diatomaceous earths, or the purified silicates. Effective surfactants, comprising from about 0.5% to about 10% of the wettable powder, are found among the sulfonated lignins, the condensed naphthalenesulfonates, the naphthalenesulfonates, the

alkylbenzenesulfonates, the alkyl sulfates, and nonionic surfactants such as ethylene oxide adducts of alkyl phenols.

Emulsifiable concentrates of the compounds comprise a convenient concentration of a compound, such as from about  
5 50 to about 500 grams per liter of liquid, equivalent to about 10% to about 50%, dissolved in an inert carrier which is either a water miscible solvent or a mixture of water-immiscible organic solvent and emulsifiers. Useful organic solvents include aromatics, especially the xylenes, and the  
10 petroleum fractions, especially the high-boiling naphthalenic and olefinic portions of petroleum such as heavy aromatic naphtha. Other organic solvents may also be used, such as the terpenic solvents including rosin derivatives, aliphatic ketones such as cyclohexanone, and  
15 complex alcohols such as 2-ethoxyethanol. Suitable emulsifiers for emulsifiable concentrates are chosen from conventional nonionic surfactants, such as those discussed above.

Aqueous suspensions comprise suspensions of water-  
20 insoluble compounds of this invention, dispersed in an aqueous vehicle at a concentration in the range from about 5% to about 50% by weight. Suspensions are prepared by finely grinding the compound, and vigorously mixing it into a vehicle comprised of water and surfactants chosen from the  
25 same types discussed above. Inert ingredients, such as inorganic salts and synthetic or natural gums, may also be added, to increase the density and viscosity of the aqueous vehicle. It is often most effective to grind and mix the compound at the same time by preparing the aqueous mixture,  
30 and homogenizing it in an implement such as a sand mill, ball mill, or piston-type homogenizer.

The compounds may also be applied as granular compositions, which are particularly useful for applications

to the soil. Granular compositions usually contain from about 0.5% to about 10% by weight of the compound, dispersed in an inert carrier which consists entirely or in large part of clay or a similar inexpensive substance. Such

5 compositions are usually prepared by dissolving the compound in a suitable solvent and applying it to a granular carrier which has been pre-formed to the appropriate particle size, in the range of from about 0.5 to 3 mm. Such compositions may also be formulated by making a dough or paste of the  
10 carrier and compound and crushing and drying to obtain the desired granular particle size.

Dusts containing the compounds are prepared simply by intimately mixing the compound in powdered form with a suitable dusty agricultural carrier, such as kaolin clay,  
15 ground volcanic rock, and the like. Dusts can suitably contain from about 1% to about 10% of the compound.

It is equally practical, when desirable for any reason, to apply the compound in the form of a solution in an appropriate organic solvent, usually a bland petroleum oil,  
20 such as the spray oils, which are widely used in agricultural chemistry.

Insecticides and acaricides are generally applied in the form of a dispersion of the active ingredient in a liquid carrier. It is conventional to refer to application  
25 rates in terms of the concentration of active ingredient in the carrier. The most widely used carrier is water.

The compounds of the invention can also be applied in the form of an aerosol composition. In such compositions the active compound is dissolved or dispersed in an inert  
30 carrier, which is a pressure-generating propellant mixture. The aerosol composition is packaged in a container from which the mixture is dispensed through an atomizing valve. Propellant mixtures comprise either low-boiling halocarbons,

which may be mixed with organic solvents, or aqueous suspensions pressurized with inert gases or gaseous hydrocarbons.

The actual amount of compound to be applied to loci of insects, mites, and aphids is not critical and can readily be determined by those skilled in the art in view of the examples above. In general, concentrations of from 10 ppm to 5000 ppm of compound are expected to provide good control. With many of the compounds, concentrations of from 100 to 1500 ppm will suffice. For field crops, such as soybeans and cotton, a suitable application rate for the compounds is about 0.5 to 1.5 lb/A, typically applied in 5-20 gal/A of spray formulation containing 1200 to 3600 ppm of compound. For citrus crops, a suitable application rate is from about 100 to 1500 gal/A spray formulation, which is a rate of 100 to 1000 ppm.

The locus to which a compound is applied can be any locus inhabited by an insect or arachnid, for example, vegetable crops, fruit and nut trees, grape vines, and ornamental plants. Inasmuch as many mite species are specific to a particular host, the foregoing list of mite species provides exemplification of the wide range of settings in which the present compounds can be used.

Because of the unique ability of mite eggs to resist toxicant action, repeated applications may be desirable to control newly emerged larvae, as is true of other known acaricides.

The following formulations of compounds of the invention are typical of compositions useful in the practice of the present invention.

A. 0.75 Emulsifiable Concentrate

Compound of formula (1)	9.38%
"TOXIMUL D" (nonionic/anionic surfactant blend)	2.50%

"TOXIMUL H" (nonionic/anionic surfactant blend)	2.50%
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"EXXON 200" (naphthalenic solvent)	85.62%
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B. 1.5 Emulsifiable Concentrate

Compound of formula (1)	18.50%
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"TOXIMUL D"	2.50%
-------------	-------

"TOXIMUL H"	2.50%
-------------	-------

"EXXON 200"	76.50%
-------------	--------

C. 1.0 Emulsifiable Concentrate

Compound of formula (1)	12.5%
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N-methylpyrrolidone	25.00%
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"TOXIMUL D"	2.50%
-------------	-------

"TOXIMUL H"	2.50%
-------------	-------

"EXXON 200"	57.50%
-------------	--------

D. 1.0 Aqueous Suspension

Compound of formula (1)	12.00%
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"PLURONIC P-103" (block copolymer of propylene oxide and ethylene oxide, surfactant)	1.50%
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"PROXEL GXL" (bio~ide/preservative)	.05%
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"AF-100" (silicon based antifoam agent)	.20%
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"REAX 88B" (lignosulfonate dispersing agent)	1.00%
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propylene glycol	10.00%
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veegum	.75%
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xanthan	.25%
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water	74.25%
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E. 1.0 Aqueous Suspension

Compound of formula (1)	12.50%
"MAKON 10" (10 moles ethyleneoxide nonylphenol surfactant)	1.00%
"ZEOSYL 200" (silica)	1.00%
"AF-100"	0.20%
"AGRIWET FR" (surfactant)	3.00%
2% xanthan hydrate	10.00%
water	72.30%

F. 1.0 Aqueous Suspension

Compound of formula (1)	12.50%
"MAKON 10"	1.50%
"ZEOSYL 200" (silica)	1.00%
"AF-100"	0.20%
"POLYFON H" (lignosulfonate dispersing agent)	0.20%
2% xanthan hydrate	10.00%
water	74.60%

G. Wettable Powder

Compound of formula (1)	25.80%
"POLYFON H"	3.50%
"SELLOGEN HR"	5.00%
"STEPANOL ME DRY"	1.00%
gum arabic	0.50%
"HISIL 233"	2.50%
Barden clay	61.70%

H. 1.0 Aqueous Suspension

Compound of formula (1)	12.40%
"TERGITOL 158-7"	5.00%
" ZEOSYL 200 "	1.0%
"AF-1G0"	0.20%
"POLYFON H"	0.50%
2% xanthan solution	10.00%
tap water	70.90%



I. 1.0 Emulsifiable Concentrate

Compound of formula (1)	12.40%
"TOXIMUL D"	2.50%
"TOXIMUL H"	2.50%
"EXXON 200"	82.60%

J. Wettable Powder

Compound of formula (1)	25.80%
"SELLOGEN HR"	5.00%
"POLYFON H"	4.00%
"STEPANOL ME DRY"	2.00%
"HISIL 233"	3.00%
Barden clay	60.20%

K. 0.5 Emulsifiable Concentrate

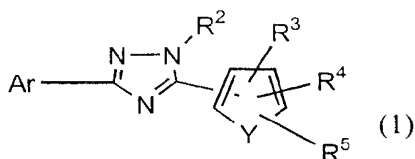
Compound of formula (1)	6.19%
"TOXIMUL H"	3.60%
"TOXIMUL D"	0.40%
"EXXON 200"	89.81%

L. Emulsifiable Concentrate

Compound of formula (1)	5 to 48
surfactant or surfactant blend	2 to 20%
Aromatic Solvent or Mixture	55 to 75%

We claim:

1. A compound of the formula (1)



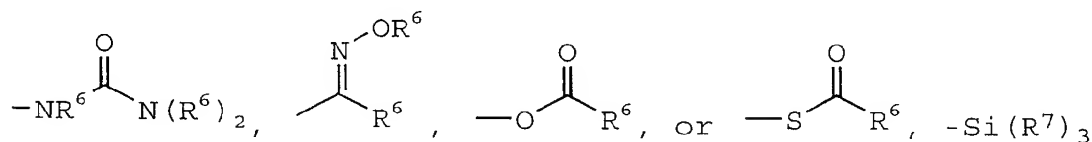
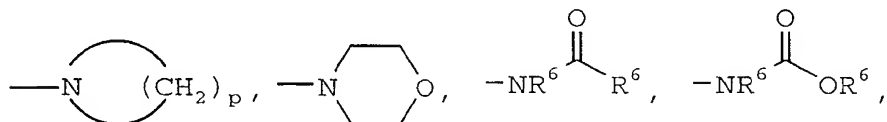
wherein

- 5 Ar is substituted phenyl;

Y is O or S;

R<sup>2</sup> is lower alkyl, haloalkyl, lower alkenyl, lower alkynyl, or alkoxyalkyl;

- R<sup>3</sup> is selected from H, halo, lower alkyl, (C<sub>7</sub>-C<sub>21</sub>)  
 10 straight or branched chain alkyl, hydroxy, lower alkoxy, haloalkyl, haloalkoxy, alkoxyalkyl, alkoxyalkoxy, lower alkenyl, lower alkynyl, haloalkenyl, CN, NO<sub>2</sub>, COR<sup>6</sup>, CO<sub>2</sub>R<sup>6</sup>, CON(R<sup>6</sup>)<sub>2</sub>, (C<sub>3</sub>-C<sub>6</sub>) cycloalkyl, S(O)<sub>m</sub>R<sup>6</sup>, -OSO<sub>2</sub>R<sup>6</sup>, SCN, -  
 (CH<sub>2</sub>)<sub>n</sub>R<sup>6</sup>, -CH=CHR<sup>6</sup>, -C≡CR<sup>6</sup>, -(CH<sub>2</sub>)<sub>q</sub>OR<sup>6</sup>, -(CH<sub>2</sub>)<sub>q</sub>SR<sup>6</sup>, -  
 15 (CH<sub>2</sub>)<sub>q</sub>NR<sup>6</sup>R<sup>6</sup>, -O(CH<sub>2</sub>)<sub>q</sub>R<sup>6</sup>, -S(CH<sub>2</sub>)<sub>q</sub>R<sup>6</sup>, -NR<sup>6</sup>(CH<sub>2</sub>)<sub>q</sub>R<sup>6</sup>,



- pyridyl, substituted pyridyl, isoxazolyl, substituted  
 isoxazolyl, naphthyl, substituted naphthyl, phenyl,  
 20 substituted phenyl, thienyl, substituted thienyl, pyrimidyl,  
 substituted pyrimidyl, pyrazolyl, or substituted pyrazolyl;

$R^4$  and  $R^5$  are independently H, halo, lower alkyl, lower alkoxy, haloalkyl, haloalkoxy, CN,  $\text{CO}_2R^6$ ,  $\text{CON}(R^6)_2$ , or  $\text{S}(\text{O})_m$  alkyl, or

if  $R^4$  and  $R^5$  are attached to adjacent carbon atoms, they may join to form a 5 or 6 member saturated or unsaturated carbocyclic ring which may be substituted by 1 or 2 halo, lower alkyl, lower alkoxy or haloalkyl groups;

$R^6$  is H, lower alkyl, haloalkyl, lower alkenyl, lower alkynyl, phenyl, or substituted phenyl;

$R^7$  is lower alkyl;

$m$  is 0, 1, or 2;

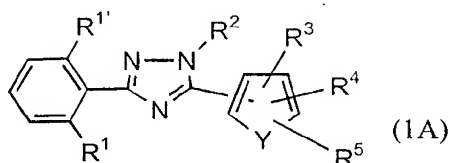
$n$  is 1 or 2;

$p$  is an integer from 2 to 6; and

$q$  is 0 or 1;

or a phytolegically acceptable acid addition salt thereof.

2. A compound of the formula (1A)



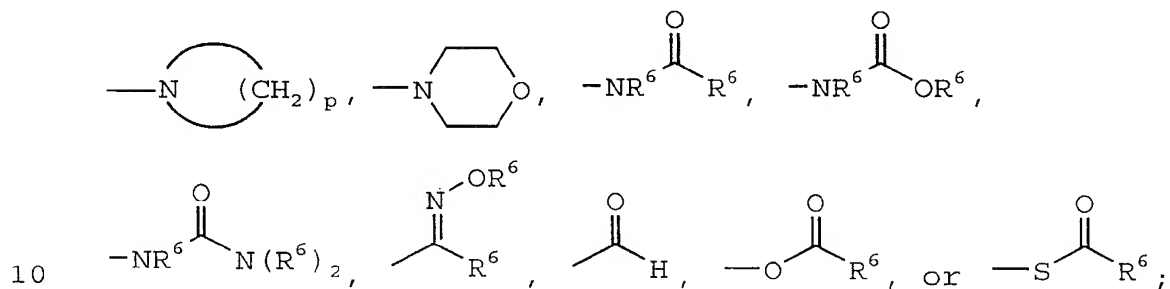
wherein

$R^1$  and  $R^{1'}$  are independently H, Cl, F, methyl, halomethyl, methoxy, or halomethoxy;

$Y$  is O or S;

$R^2$  is lower alkyl, haloalkyl, lower alkenyl, lower alkynyl, or alkoxyalkyl;

$R^3$  is selected from H, halo, lower alkyl,  $(C_7-C_{21})$  straight chain alkyl, hydroxy, lower alkoxy, haloalkyl, haloalkoxy, alkoxyalkyl, alkoxyalkoxy, lower alkenyl, haloalkenyl, CN,  $NO_2$ ,  $CO_2R^6$ ,  $CON(R^6)_2$ ,  $(C_3-C_6)$  cycloalkyl,  
 5  $S(O)_mR^6$ , SCN, pyridyl, substituted pyridyl, isoxazolyl, substituted isoxazolyl, naphthyl, substituted naphthyl, phenyl, substituted phenyl,  $-(CH_2)_nR^6$ ,  $-CH=CHR^6$ ,  $-C\equiv CR^6$ ,  $-CH_2OR^6$ ,  $-CH_2SR^6$ ,  $-CH_2NR^6R^6$ ,  $-OCH_2R^6$ ,  $-SCH_2R^6$ ,  $-NR^6CH_2R^6$ ,



$R^4$  and  $R^5$  are independently H, halo, lower alkyl, lower alkoxy, haloalkyl, haloalkoxy, CN,  $CO_2R^6$ ,  $CON(R^6)_2$ , or  $S(O)_m$  alkyl, or

$R^4$  and  $R^5$  form a 5 or 6 member saturated or unsaturated  
 15 carbocyclic ring which may be substituted by 1 or 2 halo, lower alkyl, lower alkoxy or haloalkyl groups;

$R^6$  is H, lower alkyl, haloalkyl, lower alkenyl, lower alkynyl, phenyl, or substituted phenyl;

$m$  is 0, 1, or 2; and

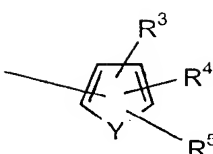
20  $n$  is 1 or 2;

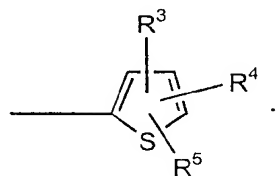
$p$  is an integer from 2 to 6;

or a phytolegically acceptable acid addition salt thereof.

3. A compound of claim 2 wherein Y is sulfur.

25 4. A compound of claim 2 wherein

the group  is a 2-thienyl group of the

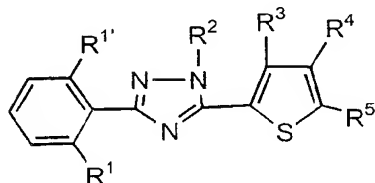
formula .

5 5. A compound of claim 2 wherein  $R^1$  is chloro and  $R^{1'}$  is chloro or fluoro.

6. A compound of claim 2 wherein  $R^2$  is methyl.

7. A compound of claim 2 wherein  $R^3$ ,  $R^4$ , and  $R^5$  are each halo.

8. A compound of formula (1B)



10

wherein

$R^1$  and  $R^{1'}$  are independently F or Cl;

$R^2$  is lower alkyl, with methyl being most preferred; and

$R^3$ ,  $R^4$  and  $R^5$  are independently H, Cl, or Br.

15 9. A compound of claim 8 wherein  $R^1$  is F or Cl and  $R^{1'}$  is F.

10. The compound of claim 9 which is 3-(2-chloro-6-fluorophenyl)-5-(3,4,5-trichlorothiophen-2-yl)-1-methyl[1,2,4]triazole.

20 11. A compound of claim 8 wherein  $R^1$  and  $R^{1'}$  are Cl.

12. The compound of claim 12 which is 3-(2,6-dichlorophenyl)-5-(3,4,5-trichlorothien-2-yl)-1-methyl[1,2,4]triazole.

5 13. A composition for controlling insects or mites which comprises a compound of claim 1 in combination with a phytologically-acceptable carrier.

10 14. A method of controlling insects or mites which comprises applying to a locus where control is desired an insect- or mite-inactivating amount of a compound of claim 1.

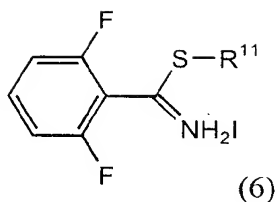
15 15. A method of controlling whitefly which comprises applying to a locus where control is desired a whitefly inactivating amount of a compound of claim 1.

20 16. A method of controlling mites which comprises applying to a locus where control is desired a mite-inactivating amount of a compound of claim 1.

25 17. A method of controlling aphids which comprises applying to a locus where control is desired an aphid inactivating amount of a compound of claim 1.

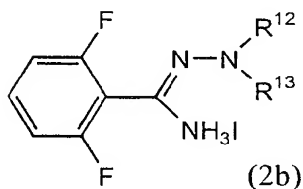
30 18. A method of protecting a plant from aphids, mites, or insects which comprises treating plant seed prior to planting it, treating soil where plant seed is to be planted, or treating soil at the roots of a plant after it is planted, with an effective amount of a compound of claim 1.

19. A compound of the formula (6)



wherein R<sup>11</sup> is lower alkyl.

20. A compound of the formula (2b)



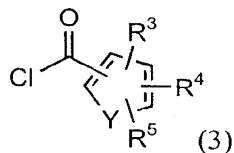
5 wherein R<sup>12</sup> is lower alkyl, and R<sup>13</sup> is a conventional amino protecting group.

21. A process for preparing a compound of claim 1 wherein R<sup>1</sup> and R<sup>1'</sup> are F which comprises reacting a compound of formula (2b)



wherein R<sup>12</sup> is lower alkyl, and R<sup>13</sup> is a conventional amino protecting group,

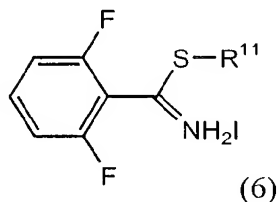
with a compound of formula (3)



15 wherein R<sup>3</sup>, R<sup>4</sup>, and R<sup>5</sup> are independently selected from H, halo, and lower alkyl.

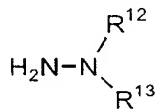
22. A process for preparing a compound of claim 1 wherein  $R^1$  and  $R^{1'}$  are F which comprises the steps of

(a) reacting a compound of formula (6)

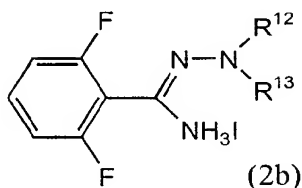


5 wherein  $R^{11}$  is lower alkyl,

with a compound of formula



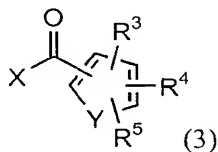
to produce a compound of formula (2b)



10 wherein  $R^{12}$  is lower alkyl, and  $R^{13}$  is a conventional amino protecting group;

and

(b) reacting the compound of formula (2b) with a compound of formula (3)



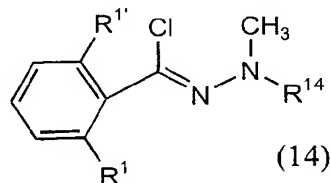
15



wherein  $R^3$ ,  $R^4$ , and  $R^5$  are independently selected from H, halo, and lower alkyl.

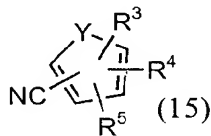
23. A process for preparing a compound of claim 1 wherein  $R^2$  is  $CH_3$  which comprises

5 reacting the compound of formula (14)



wherein  $R^1$  and  $R^{1'}$  are as defined in claim 1, and  $R^{14}$  is methanesulfonyl, benzenesulfonyl, or substituted benzenesulfonyl

10 with a compound of formula (15)

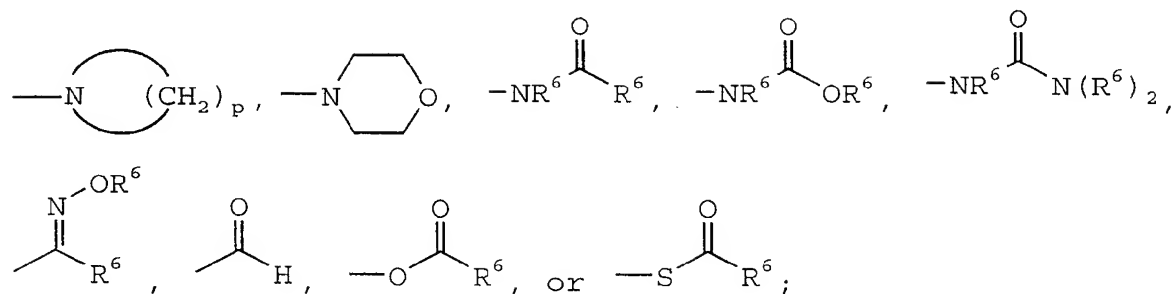


wherein

$R^3$  is selected from H, halo, lower alkyl,  $(C_7-C_{21})$  straight chain alkyl, hydroxy, lower alkoxy, haloalkyl, haloalkoxy, alkoxyalkyl, alkoxyalkoxy, lower alkenyl, haloalkenyl, CN,  $NO_2$ ,  $CO_2R^6$ ,  $CON(R^6)_2$ ,  $(C_3-C_6)$  cycloalkyl,  $S(O)_mR^6$ , SCN, pyridyl, substituted pyridyl, isoxazolyl, substituted isoxazolyl, naphthyl, substituted naphthyl, phenyl, substituted phenyl,  $-(CH_2)_nR^6$ ,  $-CH=CHR^6$ ,  $-C\equiv CR^6$ ,  $-CH_2OR^6$ ,  $-CH_2SR^6$ ,  $-CH_2NR^6R^6$ ,  $-OCH_2R^6$ ,  $-SCH_2R^6$ ,  $-NR^6CH_2R^6$ ,

15

20



R<sup>4</sup> and R<sup>5</sup> are independently H, halo, lower alkyl, lower alkoxy, haloalkyl, haloalkoxy, CN, CO<sub>2</sub>R<sup>6</sup>, CON(R<sup>6</sup>)<sub>2</sub>, or S(O)<sub>m</sub> alkyl, or

if R<sup>4</sup> and R<sup>5</sup> are attached to adjacent carbon atoms, they may form a 5 or 6 member saturated or unsaturated carbocyclic ring which may be substituted by 1 or 2 halo, lower alkyl, lower alkoxy or haloalkyl groups;

R<sup>6</sup> is H, lower alkyl, haloalkyl, lower alkenyl, lower alkynyl, phenyl, or substituted phenyl;

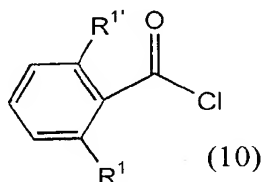
m is 0, 1, or 2; and

n is 1 or 2; and

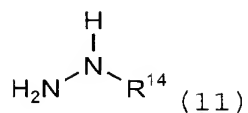
p is an integer from 2 to 6.

24. A process for preparing a compound of claim 1 wherein R<sup>2</sup> is CH<sub>3</sub> which comprises the steps of

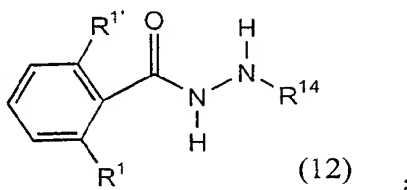
a) reacting a compound of formula (10)



wherein R<sup>1</sup> and R<sup>1'</sup> are independently H, Cl, F, methyl, halomethyl, methoxy, or halomethoxy, with a compound of formula (11)

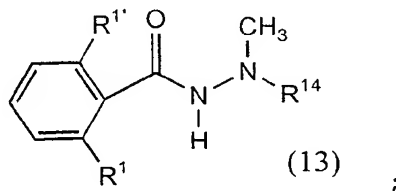


wherein  $\text{R}^{14}$  is methanesulfonyl, benzenesulfonyl, or substituted benzenesulfonyl, to produce a compound of formula (12)



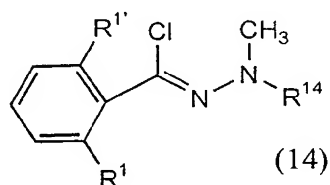
5

b) methylating the compound of formula (12) to produce a compound of formula (13)

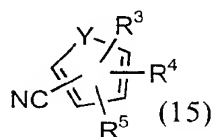


c) chlorinating the compound of formula (13)

10 to produce a compound of formula (14)

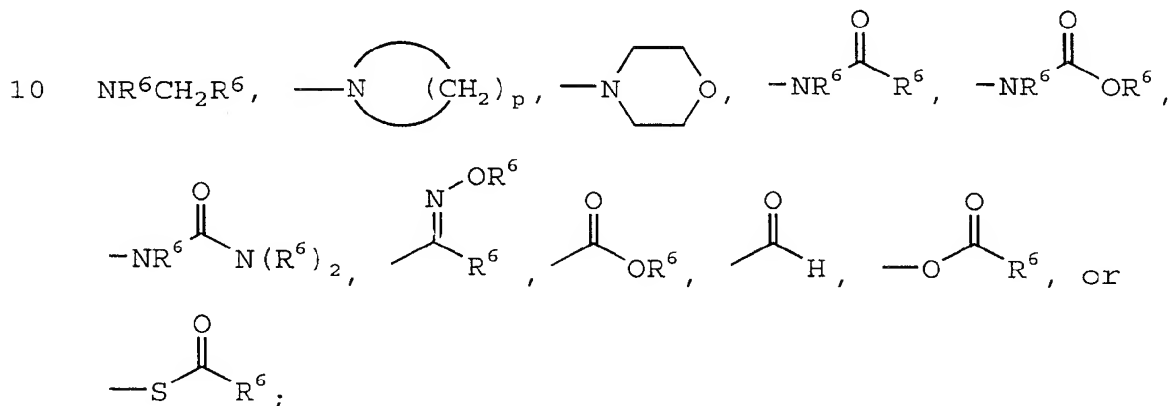


d) reacting the compound of formula (14) with a compound of formula (15)



wherein

$R^3$  is selected from H, halo, lower alkyl, ( $C_7-C_{21}$ ) straight chain alkyl, hydroxy, lower alkoxy, haloalkyl, haloalkoxy, alkoxyalkyl, alkoxyalkoxy, lower alkenyl, haloalkenyl, CN,  $NO_2$ ,  $CO_2R^6$ ,  $CONHR^6$ ,  $CON(R^6)_2$ , ( $C_3-C_6$ ) cycloalkyl,  $S(O)_mR^6$ , SCN, pyridyl, substituted pyridyl, isoxazolyl, substituted isoxazolyl, naphthyl, substituted naphthyl, phenyl, substituted phenyl,  $-(CH_2)_nR^6$ ,  $-CH=CHR^6$ ,  $-C\equiv CR^6$ ,  $-CH_2OR^6$ ,  $-CH_2SR^6$ ,  $-CH_2NR^6R^6$ ,  $-OCH_2R^6$ ,  $-SCH_2R^6$ , -



$R^4$  and  $R^5$  are independently H, halo, lower alkyl, lower alkoxy, haloalkyl, haloalkoxy, CN,  $CO_2R^6$ ,  $CONHR^6$ ,  $CON(R^6)_2$ ,  
 15 or  $S(O)_m$  alkyl, or

$R^4$  and  $R^5$  form a 5 or 6 member saturated or unsaturated carbocyclic ring which may be substituted by 1 or 2 halo, lower alkyl, lower alkoxy or haloalkyl groups;

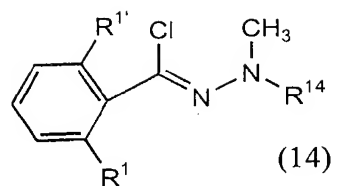
$R^6$  is H, lower alkyl, haloalkyl, lower alkenyl, lower alkynyl, phenyl, or substituted phenyl;  
 20

$m$  is 0, 1, or 2; and

$n$  is 1 or 2; and

$p$  is an integer from 2 to 6.

25. A compound of the formula (14)



wherein  $R^1$  and  $R^{1'}$  are independently H, Cl, F, methyl, halomethyl, methoxy, or halomethoxy, and

$R^{14}$  is methanesulfonyl, benzenesulfonyl, or substituted  
5 benzenesulfonyl.

# INTERNATIONAL SEARCH REPORT

International Application No

PCT/US 98/05317

## A. CLASSIFICATION OF SUBJECT MATTER

IPC 6 C07D409/04 C07D405/04 C07D409/14 C07D413/14 A01N43/653  
C07C327/58 C07C281/04 C07C257/22 C07C311/49

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 6 C07D C07C A01N

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	EP 0 572 142 A (KUMIAI CHEMICAL INDUSTRY CO., LTD ET AL.) 1 December 1993 cited in the application see the whole document ---	1-18, 21-25
X	EP 0 648 752 A (KUMIAI CHEMICAL INDUSTRY CO., LTD. ET AL.) 19 April 1995 see the whole document ---	1-18, 21-25
X	US 5 284 860 A (M. OZAKI ET AL.) 8 February 1994 cited in the application see the whole document ---	1-18, 21-25
X	US 5 380 944 A (M. OSAKI ET AL.) 10 January 1995 cited in the application see the whole document ---	1-18, 21-25
	-/--	



Further documents are listed in the continuation of box C.



Patent family members are listed in annex.

\* Special categories of cited documents :

- "A" document defining the general state of the art which is not considered to be of particular relevance
- "E" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

- "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- "&" document member of the same patent family

Date of the actual completion of the international search

24 August 1998

Date of mailing of the international search report

04.09.98

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Authorized officer

Allard, M

# INTERNATIONAL SEARCH REPORT

International Application No

PCT/US 98/05317

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>S. ITO ET AL.:  "N-Methyl-N-(phenylsulfonyl)benzo  hydrazonoyl chloride as a potential  intermediate for nitrogen-heterocycles.  Preparation of 1-methyl-3-phenyl-  1H-1,2,4-triazoles and -pyrazoles"  BULLETIN OF THE CHEMICAL SOCIETY OF JAPAN,  vol. 56, no. 2, February 1983, pages  545-8, XP002069515  TOKYO, JP  see the whole document</p>	25
Y	<p>ATKINSON M R ET AL: "Triazoles. Part III.  Mono- and di-methyl(phenyl)-1:2:4-  triazoles"  JOURNAL OF THE CHEMICAL SOCIETY,  1954, pages 3319-24, XP002075367  London, GB  see the whole document, particularly page  3322, 5th paragraph</p>	19-22
Y	<p>METZ H-J ET AL: "Zur Synthese von  1,2,4-Triazinen, IX. Synthese von  Cyclopenta'e!-1,2,4-triazinen"  CHEMISCHE BERICHTE,  vol. 115, no. 8, 3 August 1982, pages  2807-18, XP002075368  Weinheim, DE  cited in the application  see the whole document, particularly page  2809</p>	19-22
A	<p>US 4 727 142 A (FUSS A ET AL.) 23 February  1988  see the whole document, particularly  column 3, lines 52-56, column 6, lines  4-9, and example 15</p>	19
A	<p>GB 987 253 A ("SHELL" RESEARCH LIMITED) 24  March 1965  see the whole document</p>	19

# INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 98/ 05317

## Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☐ Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:
2. ☐ Claims Nos.:  
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. ☐ Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

## Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. Claims: 1-18, 21-25
2. Claims: 19, 20

1. ☒ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

### Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
- ☒ No protest accompanied the payment of additional search fees.



# INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

PCT/US 98/05317

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
EP 572142 A	01-12-1993	BR 9302097 A	07-12-1993
		CN 1079221 A, B	08-12-1993
		DE 69303864 D	05-09-1996
		DE 69303864 T	20-03-1997
		ES 2090871 T	16-10-1996
		JP 6234751 A	23-08-1994
		US 5482951 A	09-01-1996
		US 5318959 A	07-06-1994
EP 648752 A	19-04-1995	US 5616594 A	01-04-1997
		CN 1105179 A	12-07-1995
		WO 9424110 A	27-10-1994
		JP 6345608 A	20-12-1994
		US 5700824 A	23-12-1997
US 5284860 A	08-02-1994	JP 5247029 A	24-09-1993
		BR 9300732 A	08-09-1993
		CN 1078467 A	17-11-1993
		EP 0559363 A	08-09-1993
		US 5380944 A	10-01-1995
US 5380944 A	10-01-1995	JP 5247029 A	24-09-1993
		BR 9300732 A	08-09-1993
		CN 1078467 A	17-11-1993
		EP 0559363 A	08-09-1993
		US 5284860 A	08-02-1994
US 4727142 A	23-02-1988	DE 3514450 A	23-10-1986
		EP 0200051 A	05-11-1986
		JP 61249956 A	07-11-1986
GB 987253 A		BE 612252 A	
		BE 628299 A	
		CH 425330 A	
		DE 1276402 B	
		FR 86953 E	13-07-1966
		FR 1421215 A	09-03-1966
		NL 273219 A	
		US 3318681 A	09-05-1967
		US 3338913 A	29-08-1967